

(12) INTERNATIONAL APPLICATION PUBLISHED UNDER THE PATENT COOPERATION TREATY (PCT)

(19) World Intellectual Property Organization  
International Bureau



(43) International Publication Date  
31 January 2002 (31.01.2002)

PCT

(10) International Publication Number  
WO 02/07678 A2

(51) International Patent Classification<sup>7</sup>: A61K

(21) International Application Number: PCT/US01/23125

(22) International Filing Date: 23 July 2001 (23.07.2001)

(25) Filing Language: English

(26) Publication Language: English

(30) Priority Data:

60/219,619	21 July 2000 (21.07.2000)	US
60/245,157	3 November 2000 (03.11.2000)	US
60/264,319	29 January 2001 (29.01.2001)	US
60/277,270	21 March 2001 (21.03.2001)	US

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(81) Designated States (*national*): AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, UZ, VN, YU, ZA, ZW.

(84) Designated States (*regional*): ARIPO patent (GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW), Eurasian patent (AM, AZ, BY, KG, KZ, MD, RU, TJ, TM), European patent (AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR), OAPI patent (BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG).

Published:

..... without international search report and to be republished upon receipt of that report

For two-letter codes and other abbreviations, refer to the "Guidance Notes on Codes and Abbreviations" appearing at the beginning of each regular issue of the PCT Gazette.



WO 02/07678 A2

(54) Title: MU-CONOPEPTIDES

(57) Abstract: The present invention is to  $\mu$ -cono-peptides, derivatives or pharmaceutically acceptable salts thereof. The present invention is further directed to the use of this peptide, derivatives thereof and pharmaceutically acceptable salts thereof for the treatment of disorders associated with voltage-gated sodium channels. Thus, the  $\mu$ -cono-peptides or derivatives are useful as neuromuscular blocking agents, local anesthetic agents, analgesic agents and neuroprotective agents. The  $\mu$ -cono-peptides are also useful for treating neuromuscular disorders. The invention is further directed to nucleic acid sequences encoding the  $\mu$ -cono-peptides and encoding propeptides, as well as the propeptides.

TITLE OF THE INVENTION

MU-CONOPEPTIDES

CROSS-REFERENCE TO RELATED APPLICATIONS

5 [0001] The present application claims benefit under 35 USC §119(e) to U.S. provisional patent applications Serial No. 60/219,619 filed on 21 July 2000, Serial No. 60/245,157 filed on 3 November 2000, Serial No. 60/264,319 filed on 29 January 2001 and Serial No. 60/277,270 filed on 21 March 2001. Each of these applications is incorporated herein by reference.

10 [0002] This invention was made with Government support under Grant No. PO1 GM48677 awarded by the National Institute of General Medical Sciences, National Institutes of Health, Bethesda, Maryland. The United States Government has certain rights in the invention.

BACKGROUND OF THE INVENTION

15 [0003] The present invention is to  $\mu$ -conopeptides, derivatives or pharmaceutically acceptable salts thereof. The present invention is further directed to the use of this peptide, derivatives thereof and pharmaceutically acceptable salts thereof for the treatment of disorders associated with voltage-gated sodium channels. Thus, the  $\mu$ -conopeptides or derivatives are useful as neuromuscular blocking agents, local anesthetic agents, analgesic agents and  
20 neuroprotective agents. The  $\mu$ -conopeptides are also useful for treating neuromuscular disorders. The invention is further directed to nucleic acid sequences encoding the  $\mu$ -conopeptides and encoding propeptides, as well as the propeptides.

[0004] The publications and other materials used herein to illuminate the background of the invention, and in particular, cases to provide additional details respecting the practice, are  
25 incorporated by reference, and for convenience are referenced in the following text by author and date and are listed alphabetically by author in the appended bibliography.

[0005] *Conus* is a genus of predatory marine gastropods (snails) which envenomate their prey. Venomous cone snails use a highly developed projectile apparatus to deliver their cocktail of toxic conotoxins into their prey. In fish-eating species such as *Conus magus* the cone detects  
30 the presence of the fish using chemosensors in its siphon and when close enough extends its proboscis and fires a hollow harpoon-like tooth containing venom into the fish. This immobilizes the fish and enables the cone snail to wind it into its mouth via an attached filament. For general information on *Conus* and their venom see the website address

<http://grimwade.biochem.unimelb.edu.au/cone/referenc.html>. Prey capture is accomplished through a sophisticated arsenal of peptides which target specific ion channel and receptor subtypes. Each *Conus* species venom appears to contain a unique set of 50-200 peptides. The composition of the venom differs greatly between species and between individual snails within each species, each optimally evolved to paralyse its prey. The active components of the venom are small peptides toxins, typically 10-40 amino acid residues in length and are typically highly constrained peptides due to their high density of disulphide bonds.

[0006] The venoms consist of a large number of different peptide components that when separated exhibit a range of biological activities: when injected into mice they elicit a range of physiological responses from shaking to depression. The paralytic components of the venom that have been the focus of recent investigation are the  $\alpha$ -,  $\omega$ - and  $\mu$ -conotoxins. All of these conotoxins act by preventing neuronal communication, but each targets a different aspect of the process to achieve this. The  $\alpha$ -conotoxins target nicotinic ligand gated channels, the  $\mu$ -conotoxins target the voltage-gated sodium channels and the  $\omega$ -conotoxins target the voltage-gated calcium channels (Olivera et al., 1985; Olivera et al., 1990). For example a linkage has been established between  $\alpha$ -,  $\alpha$ A- &  $\phi$ -conotoxins and the nicotinic ligand-gated ion channel;  $\omega$ -conotoxins and the voltage-gated calcium channel;  $\mu$ -conotoxins and the voltage-gated sodium channel;  $\delta$ -conotoxins and the voltage-gated sodium channel;  $\kappa$ -conotoxins and the voltage-gated potassium channel; conantokins and the ligand-gated glutamate (NMDA) channel.

[0007] However, the structure and function of only a small minority of these peptides have been determined to date. For peptides where function has been determined, three classes of targets have been elucidated: voltage-gated ion channels; ligand-gated ion channels, and G-protein-linked receptors.

[0008] *Conus* peptides which target voltage-gated ion channels include those that delay the inactivation of sodium channels, as well as blockers specific for sodium channels, calcium channels and potassium channels. Peptides that target ligand-gated ion channels include antagonists of NMDA and serotonin receptors, as well as competitive and noncompetitive nicotinic receptor antagonists. Peptides which act on G-protein receptors include neurotensin and vasopressin receptor agonists. The unprecedented pharmaceutical selectivity of conotoxins is at least in part defined by a specific disulfide bond frameworks combined with hypervariable amino acids within disulfide loops (for a review see McIntosh et al., 1998).

[0009] There are drugs used in the treatment of pain, which are known in the literature and to the skilled artisan. See, for example, Merck Manual, 16th Ed. (1992). However, there is a demand for more active analgesic agents with diminished side effects and toxicity and which are non-addictive. The ideal analgesic would reduce the awareness of pain, produce analgesia over a wide range of pain types, act satisfactorily whether given orally or parenterally, produce minimal or no side effects, be free from tendency to produce tolerance and drug dependence.

[0010] Due to the high potency and exquisite selectivity of the conopeptides, several are in various stages of clinical development for treatment of human disorders. For example, two *Conus* peptides are being developed for the treatment of pain. The most advanced is  $\omega$ -conotoxin MVIIA (ziconotide), an N-type calcium channel blocker (see Heading, C., 1999; U.S. Patent No. 5,859,186).  $\omega$ -Conotoxin MVIIA, isolated from *Conus magus*, is approximately 1000 times more potent than morphine, yet does not produce the tolerance or addictive properties of opiates.  $\omega$ -Conotoxin MVIIA has completed Phase III (final stages) of human clinical trials and has been approved as a therapeutic agent.  $\omega$ -Conotoxin MVIIA is introduced into human patients by means of an implantable, programmable pump with a catheter threaded into the intrathecal space. Preclinical testing for use in post-surgical pain is being carried out on another *Conus* peptide, contulakin-G, isolated from *Conus geographus* (Craig et al. 1999). Contulakin-G is a 16 amino acid O-linked glycopeptide whose C-terminus resembles neurotensin. It is an agonist of neurotensin receptors, but appears significantly more potent than neurotensin in inhibiting pain in *in vivo* assays.

[0011] In view of a large number of biologically active substances in *Conus* species it is desirable to further characterize them and to identify peptides capable of treating disorders involving voltage gated ion channels, such as stroke and pain. Surprisingly, and in accordance with this invention, Applicants have discovered novel conotoxins that can be useful for the treatment of disorders involving voltage gated ion channels and could address a long felt need for a safe and effective treatment.

#### SUMMARY OF THE INVENTION

[0012] The present invention is to  $\mu$ -conopeptides, derivatives or pharmaceutically acceptable salts thereof. The present invention is further directed to the use of this peptide, derivatives thereof and pharmaceutically acceptable salts thereof for the treatment of disorders associated with voltage-gated sodium channels. Thus, the  $\mu$ -conopeptides or derivatives are

useful as neuromuscular blocking agents, local anesthetic agents, analgesic agents and neuroprotective agents. The  $\mu$ -conopeptides are also useful for treating neuromuscular disorders. The invention is further directed to nucleic acid sequences encoding the  $\mu$ -conopeptides and encoding propeptides, as well as the propeptides.

5 [0013] More specifically, the present invention is directed to  $\mu$ -conopeptides, having the amino acid sequences set forth in Tables 1 and 2 below.

[0014] The present invention is also directed to derivatives or pharmaceutically acceptable salts of the  $\mu$ -conopeptides or the derivatives. Examples of derivatives include peptides in which the Arg residues may be substituted by Lys, ornithine, homoargine, nor-Lys,  
10 N-methyl-Lys, N,N-dimethyl-Lys, N,N,N-trimethyl-Lys or any synthetic basic amino acid; the Lys residues may be substituted by Arg, ornithine, homoargine, nor-Lys, or any synthetic basic amino acid; the Tyr residues may be substituted with meta-Tyr, ortho-Tyr, nor-Tyr, mono-halo-Tyr, di-halo-Tyr, O-sulpho-Tyr, O-phospho-Tyr, nitro-Tyr or any synthetic hydroxy containing amino acid; the Ser residues may be substituted with Thr or any synthetic hydroxylated amino  
15 acid; the Thr residues may be substituted with Ser or any synthetic hydroxylated amino acid; the Phe residues may be substituted with any synthetic aromatic amino acid; the Trp residues may be substituted with Trp (D), neo-Trp, halo-Trp (D or L) or any aromatic synthetic amino acid; and the Asn, Ser, Thr or Hyp residues may be glycosylated. The halogen may be iodo, chloro, fluoro or bromo; preferably iodo for halogen substituted-Tyr and bromo for halogen-substituted  
20 Trp. The Tyr residues may also be substituted with the 3-hydroxyl or 2-hydroxyl isomers (meta-Tyr or ortho-Tyr, respectively) and corresponding O-sulpho- and O-phospho-derivatives. The acidic amino acid residues may be substituted with any synthetic acidic amino acid, e.g., tetrazolyl derivatives of Gly and Ala. The aliphatic amino acids may be substituted by synthetic derivatives bearing non-natural aliphatic branched or linear side chains  $C_nH_{2n+2}$  up to and  
25 including  $n=8$ . The Met residues may be substituted by norleucine (Nle). The Cys residues may be in D or L configuration and may optionally be substituted with homocysteine (D or L).

[0015] Examples of synthetic aromatic amino acid include, but are not limited to, nitro-Phe, 4-substituted-Phe wherein the substituent is  $C_1$ - $C_3$  alkyl, carboxyl, hydroxymethyl, sulphomethyl, halo, phenyl, -CHO, -CN, -SO<sub>3</sub>H and -NHAc. Examples of synthetic hydroxy  
30 containing amino acid, include, but are not limited to, such as 4-hydroxymethyl-Phe, 4-hydroxyphenyl-Gly, 2,6-dimethyl-Tyr and 5-amino-Tyr. Examples of synthetic basic amino acids include, but are not limited to, N-1-(2-pyrazoliny)-Arg, 2-(4-piperiny)-Gly, 2-(4-

piperinyl)-Ala, 2-[3-(2S)pyrrolinyl]-Gly and 2-[3-(2S)pyrrolinyl]-Ala. These and other synthetic basic amino acids, synthetic hydroxy containing amino acids or synthetic aromatic amino acids are described in Building Block Index, Version 3.0 (1999 Catalog, pages 4-47 for hydroxy containing amino acids and aromatic amino acids and pages 66-87 for basic amino acids; see also <http://www.amino-acids.com>), incorporated herein by reference, by and available from RSP Amino Acid Analogues, Inc., Worcester, MA. Examples of synthetic acid amino acids include those derivatives bearing acidic functionality, including carboxyl, phosphate, sulfonate and synthetic tetrazolyl derivatives such as described by Ornstein et al. (1993) and in U.S. Patent No. 5,331,001, each incorporated herein by reference.

[0016] Optionally, in the  $\mu$ -conopeptides of the present invention, the Asn residues may be modified to contain an N-glycan and the Ser, Thr and Hyp residues may be modified to contain an O-glycan (e.g., g-N, g-S, g-T and g-Hyp). In accordance with the present invention, a glycan shall mean any N-, S- or O-linked mono-, di-, tri-, poly- or oligosaccharide that can be attached to any hydroxy, amino or thiol group of natural or modified amino acids by synthetic or enzymatic methodologies known in the art. The monosaccharides making up the glycan can include D-allose, D-altrose, D-glucose, D-mannose, D-gulose, D-idose, D-galactose, D-talose, D-galactosamine, D-glucosamine, D-N-acetyl-glucosamine (GlcNAc), D-N-acetyl-galactosamine (GalNAc), D-fucose or D-arabinose. These saccharides may be structurally modified, e.g., with one or more O-sulfate, O-phosphate, O-acetyl or acidic groups, such as sialic acid, including combinations thereof. The glycan may also include similar polyhydroxy groups, such as D-penicillamine 2,5 and halogenated derivatives thereof or polypropylene glycol derivatives. The glycosidic linkage is beta and 1-4 or 1-3, preferably 1-3. The linkage between the glycan and the amino acid may be alpha or beta, preferably alpha and is 1-.

[0017] Core O-glycans have been described by Van de Steen et al. (1998), incorporated herein by reference. Mucin type O-linked oligosaccharides are attached to Ser or Thr (or other hydroxylated residues of the present peptides) by a GalNAc residue. The monosaccharide building blocks and the linkage attached to this first GalNAc residue define the "core glycans," of which eight have been identified. The type of glycosidic linkage (orientation and connectivities) are defined for each core glycan. Suitable glycans and glycan analogs are described further in U.S. Serial No. 09/420,797 filed 19 October 1999 and in PCT Application No. PCT/US99/24380 filed 19 October 1999 (PCT Published Application No. WO 00/23092), each incorporated herein by reference. A preferred glycan is Gal( $\beta$ 1 $\rightarrow$ 3)GalNAc( $\alpha$ 1 $\rightarrow$ ).

[0018] Optionally, in the  $\mu$ -conopeptides described above, pairs of Cys residues may be replaced pairwise with isoteric lactam or ester-thioether replacements, such as Ser/(Glu or Asp), Lys/(Glu or Asp), Cys/(Glu or Asp) or Cys/Ala combinations. Sequential coupling by known methods (Barnay et al., 2000; Hruby et al., 1994; Bitan et al., 1997) allows replacement of native  
5 Cys bridges with lactam bridges. Thioether analogs may be readily synthesized using halo-Ala residues commercially available from RSP Amino Acid Analogues.

[0019] The present invention is further directed to derivatives of the above peptides and peptide derivatives which are acyclic permutations in which the cyclic permutants retain the native bridging pattern of native toxin. See, for example, Craik et al. (2001).

10 [0020] The present invention is further directed to a method of treating disorders associated with voltage gated ion channel disorders in a subject comprising administering to the subject an effective amount of the pharmaceutical composition comprising a therapeutically effective amount of a  $\mu$ -conopeptide described herein or a pharmaceutically acceptable salt or solvate thereof. The present invention is also directed to a pharmaceutical composition  
15 comprising a therapeutically effective amount of a  $\mu$ -conopeptide described herein or a pharmaceutically acceptable salt or solvate thereof and a pharmaceutically acceptable carrier.

[0021] More specifically, the present invention is further directed to uses of these peptides or nucleic acids as described herein as neuromuscular blocking agents, local anesthetic agents, analgesic agents and neuroprotective agents. The  $\mu$ -conopeptides are also useful for  
20 treating neuromuscular disorders.

[0022] The present invention is directed to the use of  $\mu$ -conopeptides as a local anesthetic for treating pain. The  $\mu$ -conopeptides have long lasting anesthetic activity and are particularly useful for spinal anesthesia, either administered acutely for post-operative pain or via an intrathecal pump for severe chronic pain situations. The  $\mu$ -conopeptides are also useful as  
25 analgesics in chronic and neuropathic pain states, such as trigeminal neuralgia, diabetic neuropathy, post-herpetic neuralgia, neuroma pain and phantom limb pain. The  $\mu$ -conopeptides are also useful for treating burn pain and as ocular anesthetics.

[0023] The present invention is directed to the use of  $\mu$ -conopeptides as neuroprotectants. The  $\mu$ -conopeptides are useful for the treatment and alleviation of epilepsy  
30 and as a general anticonvulsant agent. The  $\mu$ -conopeptides are also useful for treating neurodegenerative diseases, such as Amyotrophic Lateral Sclerosis (ALS). The  $\mu$ -conopeptides are further useful as cerebroprotectants, such as for reducing neurotoxic injury associated with

conditions of hypoxia, anoxia or ischemia which typically follows stroke, cerebrovascular accident, brain or spinal cord trauma, myocardial infarct, physical trauma, drowning, suffocation, perinatal asphyxia, or hypoglycemic events.

[0024] The present invention is directed to the use of  $\mu$ -conopeptides as neuromuscular blockers and for treating neuromuscular disorders. As such, the  $\mu$ -conopeptides are useful for providing relaxation of muscle, for treating benign essential blepharospasm and other forms of focal dystonia and for anti-wrinkle use.

[0025] More specifically, the present invention is also directed to nucleic acids which encode  $\mu$ -conopeptides of the present invention or which encodes precursor peptides for these  $\mu$ -conopeptides, as well as the precursor peptide. The nucleic acid sequences encoding the precursor peptides of other  $\mu$ -conopeptides of the present invention are set forth in Table 1. Table 1 also sets forth the amino acid sequences of these precursor peptides.

[0026] The present invention is further directed to the use of selectively radioiodinated or radiotritiated  $\mu$ -conopeptides for characterizing pore occlusion sites on different sodium channel subtypes or for use in screening assays.

[0027] The present invention is also directed to the use of  $\mu$ -conopeptides for screening small molecule libraries to identify small molecules that are selective blocking agents at specific sodium channel subtypes expressed in mammalian systems. In one embodiment, the blocking activity of a small molecule at a particular sodium channel subtype is compared to the blocking activity of a  $\mu$ -conopeptide at the same sodium channel subtype. In a second embodiment, the ability of a small molecule to displace a  $\mu$ -conopeptide from a sodium channel subtype is determined. In a third embodiment, the binding affinity of a small molecule for a sodium channel subtype is compared to the binding affinity of a  $\mu$ -conopeptide for the same sodium channel subtype.

#### DETAILED DESCRIPTION OF THE PREFERRED EMBODIMENT

[0028] The present invention is to  $\mu$ -conopeptides, derivatives or pharmaceutically acceptable salts thereof. The present invention is further directed to the use of this peptide, derivatives thereof and pharmaceutically acceptable salts thereof for the treatment of disorders associated with voltage-gated sodium channels. Thus, the  $\mu$ -conopeptides or derivatives are useful as neuromuscular blocking agents, local anesthetic agents, analgesic agents and neuroprotective agents. The  $\mu$ -conopeptides are also useful for treating neuromuscular



disorders. The invention is further directed to nucleic acid sequences encoding the  $\mu$ -conopeptides and encoding propeptides, as well as the propeptides.

[0029] The present invention, in another aspect, relates to a pharmaceutical composition comprising an effective amount of an  $\mu$ -conopeptides, a mutein thereof, an analog thereof, an active fragment thereof or pharmaceutically acceptable salts or solvates. Such a pharmaceutical composition has the capability of acting at voltage gated ion channels, and are thus useful for treating a disorder or disease of a living animal body, including a human, which disorder or disease is responsive to the partial or complete blockade of voltage gated ion channels of the central nervous system comprising the step of administering to such a living animal body, including a human, in need thereof a therapeutically effective amount of a pharmaceutical composition of the present invention.

[0030] The present invention is directed to the use of  $\mu$ -conopeptides as neuromuscular blockers and for treating neuromuscular disorders. As such, the  $\mu$ -conopeptides are useful for providing relaxation of muscle, for treating benign essential blepharospasm and other forms of focal dystonia and for anti-wrinkle use. Thus, in one aspect, the  $\mu$ -conopeptides are useful as neuromuscular blocking agents in conjunction with surgery or for intubation of the trachea by conventional parenteral administration e.g., intramuscular or intravenous administration in solution. In a second aspect, the  $\mu$ -conopeptides are useful as agents for treating neuromuscular disorders such as myofascial pain syndrome, chronic muscle spasm, dystonias and spasticity.

[0031] The primary factor detrimental to neurons in neurological disorders associated with deficient oxygen supply or mitochondrial dysfunction is insufficient ATP production relative to their requirement. As a large part of the energy consumed by brain cells is used for maintenance of the  $\text{Na}^+$  gradient across the cellular membrane, reduction of energy demand by down-modulation of voltage-gated  $\text{Na}(+)$ -channels is one strategy for neuroprotection. In addition, preservation of the inward  $\text{Na}^+$  gradient may be beneficial because it is an essential driving force for vital ion exchanges and transport mechanisms such as  $\text{Ca}^{2+}$  homeostasis and neurotransmitter uptake. Thus, the  $\mu$ -conopeptides of the present invention are useful as neuroprotectants.

[0032] Thus, the pharmaceutical compositions of the present invention are useful as neuroprotectants, especially cerebroprotectants, neuromuscular blockers, analgesics (both as a local anesthetic and for general analgesia use) or adjuvants to general anesthetics. A "neurological disorder or disease" is a disorder or disease of the nervous system including, but

not limited to, global and focal ischemic and hemorrhagic stroke, head trauma, spinal cord injury, hypoxia-induced nerve cell damage as in cardiac arrest or neonatal distress or epilepsy. In addition, a "neurological disorder or disease" is a disease state and condition in which a neuroprotectant, anticonvulsant, analgesic and/or as an adjunct in general anesthesia may be indicated, useful, recommended or prescribed.

[0033] More specifically, the present invention is directed to the use of these compounds for reducing neurotoxic injury associated with conditions of hypoxia, anoxia or ischemia which typically follows stroke, cerebrovascular accident, brain or spinal cord trauma, myocardial infarct, physical trauma, drowning, suffocation, perinatal asphyxia, or hypoglycemic events.

The present invention is further directed to the use of these compounds for treating pain, including acute and chronic pain, such migraine, nociceptive and neuropathic pain.

[0034] A "neuroprotectant" is a compound capable of preventing the neuronal death associated with a neurological disorder or disease. An "analgesic" is a compound capable of relieving pain by altering perception of nociceptive stimuli without producing anesthesia or loss of consciousness. A "muscle relaxant" is a compound that reduces muscular tension. An "adjunct in general anesthesia" is a compound useful in conjunction with anesthetic agents in producing the loss of ability to perceive pain associated with the loss of consciousness.

[0035] The invention relates as well to methods useful for treatment of neurological disorders and diseases, including, but not limited to, global and focal ischemic and hemorrhagic stroke, head trauma, spinal cord injury, hypoxia-induced nerve cell damage such as in cardiac arrest or neonatal distress, epilepsy or other convulsive disorders without undesirable side effects.

[0036] Thus, in one embodiment, the invention provides a method of reducing/alleviating/ decreasing the perception of pain by a subject or for inducing analgesia in a subject comprising administering to the subject an effective amount of the pharmaceutical composition comprising a therapeutically effective amount of a  $\mu$ -conopeptide described herein or a pharmaceutically acceptable salt or solvate thereof. The pain may be acute, persistent, inflammatory or neuropathic pain. The  $\mu$ -conopeptides are useful as an analgesia for chronic and neuropathic pain states, such as trigeminal neuralgia, diabetic neuropathy, post-herpetic neuralgia, neuroma pain, phantom limb pain. These peptides are also useful for treating burn pain and as ocular anesthetics.

[0037] In a second embodiment, the invention provides a method of reducing/alleviating/decreasing the perception of pain by a subject or for inducing analgesia, particularly local analgesia, in a subject comprising administering to the subject an effective amount of the pharmaceutical composition comprising a therapeutically effective amount of a  $\mu$ -conopeptide described herein or a pharmaceutically acceptable salt or solvate thereof. These peptides are also  
5 useful for treating burn pain and as ocular anesthetics.

[0038] In a third embodiment, the invention provides a method of treating stroke, head or spinal cord trauma or injury, anoxia, hypoxia-induced nerve cell damage, ischemia, migraine, psychosis, anxiety, schizophrenia, inflammation, movement disorder, epilepsy, any other  
10 convulsive disorder or in the prevention of the degenerative changes connected with the same in a subject comprising administering to the subject an effective amount of the pharmaceutical composition comprising a therapeutically effective amount of a  $\mu$ -conopeptide described herein or a pharmaceutically acceptable salt or solvate thereof.

[0039] In a fourth embodiment, the invention provides a method for providing a  
15 neuromuscular block or for treating neuromuscular disorders, such as methods for providing relaxation of muscle, for treating benign essential blepharospasm and other forms of focal dystonia and for anti-wrinkle use. Thus, in one aspect, the  $\mu$ -conopeptides are useful as neuromuscular blocking agents in conjunction with surgery or for intubation of the trachea by conventional parenteral administration e.g., intramuscular or intravenous administration in  
20 solution. In a second aspect, the  $\mu$ -conopeptides are useful as agents for treating neuromuscular disorders such as myofascial pain syndrome, chronic muscle spasm, dystonias and spasticity.

[0040] The present invention is also directed to the use of  $\mu$ -conopeptides for screening small molecule libraries to identify small molecules that are selective blocking agents at specific sodium channel subtypes expressed in mammalian systems. In one embodiment, the blocking  
25 activity of a small molecule at a particular sodium channel subtype is compared to the blocking activity of a  $\mu$ -conopeptide at the same sodium channel subtype. In a second embodiment, the ability of a small molecule to displace a  $\mu$ -conopeptide from a sodium channel subtype is determined. In a third embodiment, the binding affinity of a small molecule for a sodium channel subtype is compared to the binding affinity of a  $\mu$ -conopeptide for the same sodium channel  
30 subtype.

[0041] The  $\mu$ -conopeptides described herein are sufficiently small to be chemically synthesized. General chemical syntheses for preparing the foregoing  $\omega$ -conotoxin peptides are

described hereinafter. Various ones of the  $\mu$ -conopeptides can also be obtained by isolation and purification from specific *Conus* species using the technique described in U.S. Patent Nos. 4,447,356 (Olivera et al., 1984); 5,514,774; 5,719,264; and 5,591,821, as well as in PCT published application WO 98/03189, the disclosures of which are incorporated herein by  
5 reference.

[0042] Although the  $\mu$ -conopeptides of the present invention can be obtained by purification from cone snails, because the amounts of  $\mu$ -conopeptides obtainable from individual snails are very small, the desired substantially pure  $\mu$ -conopeptides are best practically obtained in commercially valuable amounts by chemical synthesis using solid-phase strategy. For  
10 example, the yield from a single cone snail may be about 10 micrograms or less of  $\mu$ -conopeptides peptide. By "substantially pure" is meant that the peptide is present in the substantial absence of other biological molecules of the same type; it is preferably present in an amount of at least about 85% purity and preferably at least about 95% purity. Chemical synthesis of biologically active  $\mu$ -conopeptides peptides depends of course upon correct  
15 determination of the amino acid sequence.

[0043] The  $\mu$ -conopeptides can also be produced by recombinant DNA techniques well known in the art. Such techniques are described by Sambrook et al. (1989). A gene of interest (i.e., a gene that encodes a suitable  $\mu$ -conopeptides) can be inserted into a cloning site of a suitable expression vector by using standard techniques. These techniques are well known to  
20 those skilled in the art. The expression vector containing the gene of interest may then be used to transfect the desired cell line. Standard transfection techniques such as calcium phosphate co-precipitation, DEAE-dextran transfection or electroporation may be utilized. A wide variety of host/expression vector combinations may be used to express a gene encoding a conotoxin peptide of interest. Such combinations are well known to a skilled artisan. The peptides  
25 produced in this manner are isolated, reduced if necessary, and oxidized to form the correct disulfide bonds.

[0044] One method of forming disulfide bonds in the  $\mu$ -conopeptides of the present invention is the air oxidation of the linear peptides for prolonged periods under cold room temperatures or at room temperature. This procedure results in the creation of a substantial  
30 amount of the bioactive, disulfide-linked peptides. The oxidized peptides are fractionated using reverse-phase high performance liquid chromatography (HPLC) or the like, to separate peptides having different linked configurations. Thereafter, either by comparing these fractions with the

elution of the native material or by using a simple assay, the particular fraction having the correct linkage for maximum biological potency is easily determined. However, because of the dilution resulting from the presence of other fractions of less biopotency, a somewhat higher dosage may be required.

5        [0045] The peptides are synthesized by a suitable method, such as by exclusively solid-phase techniques, by partial solid-phase techniques, by fragment condensation or by classical solution couplings.

         [0046] In conventional solution phase peptide synthesis, the peptide chain can be prepared by a series of coupling reactions in which constituent amino acids are added to the  
10        growing peptide chain in the desired sequence. Use of various coupling reagents, e.g., dicyclohexylcarbodiimide or diisopropylcarbonyldimidazole, various active esters, e.g., esters of N-hydroxyphthalimide or N-hydroxy-succinimide, and the various cleavage reagents, to carry out reaction in solution, with subsequent isolation and purification of intermediates, is well known classical peptide methodology. Classical solution synthesis is described in detail in the  
15        treatise, "Methoden der Organischen Chemie (Houben-Weyl): Synthese von Peptiden," (1974). Techniques of exclusively solid-phase synthesis are set forth in the textbook, "Solid-Phase Peptide Synthesis," (Stewart and Young, 1969), and are exemplified by the disclosure of U.S. Patent 4,105,603 (Vale et al., 1978). The fragment condensation method of synthesis is exemplified in U.S. Patent 3,972,859 (1976). Other available syntheses are exemplified by U.S.  
20        Patents No. 3,842,067 (1974) and 3,862,925 (1975). The synthesis of peptides containing  $\gamma$ -carboxyglutamic acid residues is exemplified by Rivier et al. (1987), Nishiuchi et al. (1993) and Zhou et al. (1996).

         [0047] Common to such chemical syntheses is the protection of the labile side chain groups of the various amino acid moieties with suitable protecting groups which will prevent a  
25        chemical reaction from occurring at that site until the group is ultimately removed. Usually also common is the protection of an  $\alpha$ -amino group on an amino acid or a fragment while that entity reacts at the carboxyl group, followed by the selective removal of the  $\alpha$ -amino protecting group to allow subsequent reaction to take place at that location. Accordingly, it is common that, as a step in such a synthesis, an intermediate compound is produced which includes each of the  
30        amino acid residues located in its desired sequence in the peptide chain with appropriate side-chain protecting groups linked to various ones of the residues having labile side chains.

[0048] As far as the selection of a side chain amino protecting group is concerned, generally one is chosen which is not removed during deprotection of the  $\alpha$ -amino groups during the synthesis. However, for some amino acids, e.g., His, protection is not generally necessary. In selecting a particular side chain protecting group to be used in the synthesis of the peptides, the following general rules are followed: (a) the protecting group preferably retains its protecting properties and is not split off under coupling conditions, (b) the protecting group should be stable under the reaction conditions selected for removing the  $\alpha$ -amino protecting group at each step of the synthesis, and (c) the side chain protecting group must be removable, upon the completion of the synthesis containing the desired amino acid sequence, under reaction conditions that will not undesirably alter the peptide chain.

[0049] It should be possible to prepare many, or even all, of these peptides using recombinant DNA technology. However, when peptides are not so prepared, they are preferably prepared using the Merrifield solid-phase synthesis, although other equivalent chemical syntheses known in the art can also be used as previously mentioned. Solid-phase synthesis is commenced from the C-terminus of the peptide by coupling a protected  $\alpha$ -amino acid to a suitable resin. Such a starting material can be prepared by attaching an  $\alpha$ -amino-protected amino acid by an ester linkage to a chloromethylated resin or a hydroxymethyl resin, or by an amide bond to a benzhydrylamine (BHA) resin or paramethylbenzhydrylamine (MBHA) resin. Preparation of the hydroxymethyl resin is described by Bodansky et al. (1966). Chloromethylated resins are commercially available from Bio Rad Laboratories (Richmond, CA) and from Lab. Systems, Inc. The preparation of such a resin is described by Stewart and Young (1969). BHA and MBHA resin supports are commercially available, and are generally used when the desired polypeptide being synthesized has an unsubstituted amide at the C-terminus. Thus, solid resin supports may be any of those known in the art, such as one having the formulae  $-O-CH_2$ -resin support,  $-NH$  BHA resin support, or  $-NH$ -MBHA resin support. When the unsubstituted amide is desired, use of a BHA or MBHA resin is preferred, because cleavage directly gives the amide. In case the N-methyl amide is desired, it can be generated from an N-methyl BHA resin. Should other substituted amides be desired, the teaching of U.S. Patent No. 4,569,967 (Kornreich et al., 1986) can be used, or should still other groups than the free acid be desired at the C-terminus, it may be preferable to synthesize the peptide using classical methods as set forth in the Houben-Weyl text (1974).

[0050] The C-terminal amino acid, protected by Boc or Fmoc and by a side-chain protecting group, if appropriate, can be first coupled to a chloromethylated resin according to the procedure set forth in K. Horiki et al. (1978), using KF in DMF at about 60°C for 24 hours with stirring, when a peptide having free acid at the C-terminus is to be synthesized. Following the coupling of the BOC-protected amino acid to the resin support, the  $\alpha$ -amino protecting group is removed, as by using trifluoroacetic acid (TFA) in methylene chloride or TFA alone. The deprotection is carried out at a temperature between about 0°C and room temperature. Other standard cleaving reagents, such as HCl in dioxane, and conditions for removal of specific  $\alpha$ -amino protecting groups may be used as described in Schroder & Lubke (1965).

[0051] After removal of the  $\alpha$ -amino-protecting group, the remaining  $\alpha$ -amino- and side chain-protected amino acids are coupled step-wise in the desired order to obtain the intermediate compound defined hereinbefore, or as an alternative to adding each amino acid separately in the synthesis, some of them may be coupled to one another prior to addition to the solid phase reactor. Selection of an appropriate coupling reagent is within the skill of the art. Particularly suitable as a coupling reagent is N,N'-dicyclohexylcarbodiimide (DCC, DIC, HBTU, HATU, TBTU in the presence of HoBt or HoAt).

[0052] The activating reagents used in the solid phase synthesis of the peptides are well known in the peptide art. Examples of suitable activating reagents are carbodiimides, such as N,N'-diisopropylcarbodiimide and N-ethyl-N'-(3-dimethylaminopropyl)carbodiimide. Other activating reagents and their use in peptide coupling are described by Schroder & Lubke (1965) and Kapoor (1970).

[0053] Each protected amino acid or amino acid sequence is introduced into the solid-phase reactor in about a twofold or more excess, and the coupling may be carried out in a medium of dimethylformamide (DMF):CH<sub>2</sub>Cl<sub>2</sub> (1:1) or in DMF or CH<sub>2</sub>Cl<sub>2</sub> alone. In cases where intermediate coupling occurs, the coupling procedure is repeated before removal of the  $\alpha$ -amino protecting group prior to the coupling of the next amino acid. The success of the coupling reaction at each stage of the synthesis, if performed manually, is preferably monitored by the ninhydrin reaction, as described by Kaiser et al. (1970). Coupling reactions can be performed automatically, as on a Beckman 990 automatic synthesizer, using a program such as that reported in Rivier et al. (1978).

[0054] After the desired amino acid sequence has been completed, the intermediate peptide can be removed from the resin support by treatment with a reagent, such as liquid

hydrogen fluoride or TFA (if using Fmoc chemistry), which not only cleaves the peptide from the resin but also cleaves all remaining side chain protecting groups and also the  $\alpha$ -amino protecting group at the N-terminus if it was not previously removed to obtain the peptide in the form of the free acid. If Met is present in the sequence, the Boc protecting group is preferably first removed using trifluoroacetic acid (TFA)/ethanedithiol prior to cleaving the peptide from the resin with HF to eliminate potential S-alkylation. When using hydrogen fluoride or TFA for cleaving, one or more scavengers such as anisole, cresol, dimethyl sulfide and methylethyl sulfide are included in the reaction vessel.

[0055] Cyclization of the linear peptide is preferably affected, as opposed to cyclizing the peptide while a part of the peptido-resin, to create bonds between Cys residues. To effect such a disulfide cyclizing linkage, fully protected peptide can be cleaved from a hydroxymethylated resin or a chloromethylated resin support by ammonolysis, as is well known in the art, to yield the fully protected amide intermediate, which is thereafter suitably cyclized and deprotected. Alternatively, deprotection, as well as cleavage of the peptide from the above resins or a benzhydrylamine (BHA) resin or a methylbenzhydrylamine (MBHA), can take place at 0°C with hydrofluoric acid (HF) or TFA, followed by oxidation as described above.

[0056] The peptides are also synthesized using an automatic synthesizer. Amino acids are sequentially coupled to an MBHA Rink resin (typically 100 mg of resin) beginning at the C-terminus using an Advanced Chemtech 357 Automatic Peptide Synthesizer. Couplings are carried out using 1,3-diisopropylcarbodiimide in N-methylpyrrolidinone (NMP) or by 2-(1H-benzotriazole-1-yl)-1,1,3,3-tetramethyluronium hexafluorophosphate (HBTU) and diethylisopropylethylamine (DIEA). The FMOC protecting group is removed by treatment with a 20% solution of piperidine in dimethylformamide (DMF). Resins are subsequently washed with DMF (twice), followed by methanol and NMP.

[0057] Muteins, analogs or active fragments, of the foregoing conotoxin peptides are also contemplated here. See, e.g., Hammerland et al. (1992). Derivative muteins, analogs or active fragments of the conotoxin peptides may be synthesized according to known techniques, including conservative amino acid substitutions, such as outlined in U.S. Patent Nos. 5,545,723 (see particularly col. 2, line 50--col. 3, line 8); 5,534,615 (see particularly col. 19, line 45--col. 22, line 33); and 5,364,769 (see particularly col. 4, line 55--col. 7, line 26), each herein incorporated by reference.



[0058] The  $\mu$ -conopeptides of the present invention are also useful to reduce neurotoxic injury associated with conditions of hypoxia, anoxia or ischemia which typically follows stroke, cerebrovascular accident, brain or spinal chord trauma, myocardial infarct, physical trauma, drownings, suffocation, perinatal asphyxia, or hypoglycemic events. To reduce neurotoxic injury, an  $\omega$ -conopeptide should be administered in a therapeutically effective amount to the patient within 24 hours of the onset of the hypoxic, anoxic or ischemic condition in order for the  $\mu$ -conopeptide to effectively minimize the CNS damage which the patient will experience.

[0059] The  $\mu$ -conopeptides of the present invention are further useful in controlling pain, e.g., as analgesic agents, and the treatment of migraine, acute pain or persistent pain. They can be used prophylactically or to relieve the symptoms associated with a migraine episode, or to treat acute or persistent pain. For these uses, an  $\mu$ -conopeptide is administered in a therapeutically effective amount to overcome or to ease the pain.

[0060] The  $\mu$ -conopeptides of the present invention are also useful as neuromuscular blockers and for treating neuromuscular disorders. They can be used for providing relaxation of muscle, for treating benign essential blepharospasm and other forms of focal dystonia and for anti-wrinkle use. Thus, in one aspect, the  $\mu$ -conopeptides are used as neuromuscular blocking agents in conjunction with surgery or for intubation of the trachea by conventional parenteral administration e.g., intramuscular or intravenous administration in solution. In a second aspect, the  $\mu$ -conopeptides are used as agents for treating neuromuscular disorders such as myofascial pain syndrome, chronic muscle spasm, dystonias and spasticity. For these uses, a  $\mu$ -conopeptide is administered in a therapeutically effective amount to relax muscle or provide a neuromuscular block.

[0061] Pharmaceutical compositions containing a compound of the present invention as the active ingredient can be prepared according to conventional pharmaceutical compounding techniques. See, for example, *Remington's Pharmaceutical Sciences*, 18th Ed. (1990, Mack Publishing Co., Easton, PA). Typically, an antagonistic amount of active ingredient will be admixed with a pharmaceutically acceptable carrier. The carrier may take a wide variety of forms depending on the form of preparation desired for administration, e.g., intravenous, oral, parenteral or intrathecally. For examples of delivery methods see U.S. Patent No. 5,844,077, incorporated herein by reference.

[0062] "Pharmaceutical composition" means physically discrete coherent portions suitable for medical administration. "Pharmaceutical composition in dosage unit form" means

physically discrete coherent units suitable for medical administration, each containing a daily dose or a multiple (up to four times) or a sub-multiple (down to a fortieth) of a daily dose of the active compound in association with a carrier and/or enclosed within an envelope. Whether the composition contains a daily dose, or for example, a half, a third or a quarter of a daily dose, will  
5 depend on whether the pharmaceutical composition is to be administered once or, for example, twice, three times or four times a day, respectively.

[0063] The term "salt", as used herein, denotes acidic and/or basic salts, formed with inorganic or organic acids and/or bases, preferably basic salts. While pharmaceutically acceptable salts are preferred, particularly when employing the compounds of the invention as  
10 medicaments, other salts find utility, for example, in processing these compounds, or where non-medicament-type uses are contemplated. Salts of these compounds may be prepared by art-recognized techniques.

[0064] Examples of such pharmaceutically acceptable salts include, but are not limited to, inorganic and organic addition salts, such as hydrochloride, sulphates, nitrates or phosphates  
15 and acetates, trifluoroacetates, propionates, succinates, benzoates, citrates, tartrates, fumarates, maleates, methane-sulfonates, isothionates, theophylline acetates, salicylates, respectively, or the like. Lower alkyl quaternary ammonium salts and the like are suitable, as well.

[0065] As used herein, the term "pharmaceutically acceptable" carrier means a non-toxic, inert solid, semi-solid liquid filler, diluent, encapsulating material, formulation auxiliary of any  
20 type, or simply a sterile aqueous medium, such as saline. Some examples of the materials that can serve as pharmaceutically acceptable carriers are sugars, such as lactose, glucose and sucrose, starches such as corn starch and potato starch, cellulose and its derivatives such as sodium carboxymethyl cellulose, ethyl cellulose and cellulose acetate; powdered tragacanth; malt, gelatin, talc; excipients such as cocoa butter and suppository waxes; oils such as peanut oil,  
25 cottonseed oil, safflower oil, sesame oil, olive oil, corn oil and soybean oil; glycols, such as propylene glycol, polyols such as glycerin, sorbitol, mannitol and polyethylene glycol; esters such as ethyl oleate and ethyl laurate, agar; buffering agents such as magnesium hydroxide and aluminum hydroxide; alginic acid; pyrogen-free water; isotonic saline, Ringer's solution; ethyl alcohol and phosphate buffer solutions, as well as other non-toxic compatible substances used in  
30 pharmaceutical formulations.

[0066] Wetting agents, emulsifiers and lubricants such as sodium lauryl sulfate and magnesium stearate, as well as coloring agents, releasing agents, coating agents, sweetening,

flavoring and perfuming agents, preservatives and antioxidants can also be present in the composition, according to the judgment of the formulator. Examples of pharmaceutically acceptable antioxidants include, but are not limited to, water soluble antioxidants such as ascorbic acid, cysteine hydrochloride, sodium bisulfite, sodium metabisulfite, sodium sulfite, and the like; oil soluble antioxidants, such as ascorbyl palmitate, butylated hydroxyanisole (BHA), butylated hydroxytoluene (BHT), lecithin, propyl gallate, alpha-tocopherol and the like; and the metal chelating agents such as citric acid, ethylenediamine tetraacetic acid (EDTA), sorbitol, tartaric acid, phosphoric acid and the like.

[0067] For oral administration, the compounds can be formulated into solid or liquid preparations such as capsules, pills, tablets, lozenges, melts, powders, suspensions or emulsions.

In preparing the compositions in oral dosage form, any of the usual pharmaceutical media may be employed, such as, for example, water, glycols, oils, alcohols, flavoring agents, preservatives, coloring agents, suspending agents, and the like in the case of oral liquid preparations (such as, for example, suspensions, elixirs and solutions); or carriers such as starches, sugars, diluents, granulating agents, lubricants, binders, disintegrating agents and the like in the case of oral solid preparations (such as, for example, powders, capsules and tablets). Because of their ease in administration, tablets and capsules represent the most advantageous oral dosage unit form, in which case solid pharmaceutical carriers are obviously employed. If desired, tablets may be sugar-coated or enteric-coated by standard techniques. The active agent can be encapsulated to make it stable to passage through the gastrointestinal tract while at the same time allowing for passage across the blood brain barrier. See for example, WO 96/11698.

[0068] For parenteral administration, the compound may be dissolved in a pharmaceutical carrier and administered as either a solution or a suspension. Illustrative of suitable carriers are water, saline, dextrose solutions, fructose solutions, ethanol, or oils of animal, vegetative or synthetic origin. The carrier may also contain other ingredients, for example, preservatives, suspending agents, solubilizing agents, buffers and the like. When the compounds are being administered intrathecally, they may also be dissolved in cerebrospinal fluid.

[0069] A variety of administration routes are available. The particular mode selected will depend of course, upon the particular drug selected, the severity of the disease state being treated and the dosage required for therapeutic efficacy. The methods of this invention, generally speaking, may be practiced using any mode of administration that is medically acceptable,

meaning any mode that produces effective levels of the active compounds without causing clinically unacceptable adverse effects. Such modes of administration include oral, rectal, sublingual, topical, nasal, transdermal or parenteral routes. The term "parenteral" includes subcutaneous, intravenous, epidural, irrigation, intramuscular, release pumps, or infusion.

5 [0070] For example, administration of the active agent according to this invention may be achieved using any suitable delivery means, including:

(a) pump (see, e.g., Luer & Hatton (1993), Zimm et al. (1984) and Ettinger et al. (1978));

(b), microencapsulation (see, e.g., U.S. Patent Nos. 4,352,883; 4,353,888; and 5,084,350);

10 (c) continuous release polymer implants (see, e.g., U.S. Patent No. 4,883,666);

(d) macroencapsulation (see, e.g., U.S. Patent Nos. 5,284,761, 5,158,881, 4,976,859 and 4,968,733 and published PCT patent applications WO92/19195, WO 95/05452);

(e) naked or unencapsulated cell grafts to the CNS (see, e.g., U.S. Patent Nos. 5,082,670 and 5,618,531);

15 (f) injection, either subcutaneously, intravenously, intra-arterially, intramuscularly, or to other suitable site; or

(g) oral administration, in capsule, liquid, tablet, pill, or prolonged release formulation.

[0071] In one embodiment of this invention, an active agent is delivered directly into the CNS, preferably to the brain ventricles, brain parenchyma, the intrathecal space or other suitable  
20 CNS location, most preferably intrathecally.

[0072] Alternatively, targeting therapies may be used to deliver the active agent more specifically to certain types of cell, by the use of targeting systems such as antibodies or cell specific ligands. Targeting may be desirable for a variety of reasons, e.g. if the agent is unacceptably toxic, or if it would otherwise require too high a dosage, or if it would not  
25 otherwise be able to enter the target cells.

[0073] The active agents, which are peptides, can also be administered in a cell based delivery system in which a DNA sequence encoding an active agent is introduced into cells designed for implantation in the body of the patient, especially in the spinal cord region. Suitable delivery systems are described in U.S. Patent No. 5,550,050 and published PCT  
30 Application Nos. WO 92/19195, WO 94/25503, WO 95/01203, WO 95/05452, WO 96/02286, WO 96/02646, WO 96/40871, WO 96/40959 and WO 97/12635. Suitable DNA sequences can

be prepared synthetically for each active agent on the basis of the developed sequences and the known genetic code.

[0074] Exemplary methods for administering such muscle relaxant compounds (e.g., so as to achieve sterile or aseptic conditions) will be apparent to the skilled artisan. Certain methods suitable for administering compounds useful according to the present invention are set forth in Goodman and Gilman's *The Pharmacological Basis of Therapeutics*, 7th Ed. (1985). The administration to the patient can be intermittent; or at a gradual, continuous, constant or controlled rate. Administration can be to a warm-blooded animal (e.g. a mammal, such as a mouse, rat, cat, rabbit, dog, pig, cow or monkey); but advantageously is administered to a human being. Administration occurs after general anesthesia is administered. The frequency of administration normally is determined by an anesthesiologist, and typically varies from patient to patient.

[0075] The active agent is preferably administered in an therapeutically effective amount. By a "therapeutically effective amount" or simply "effective amount" of an active compound is meant a sufficient amount of the compound to treat the desired condition at a reasonable benefit/risk ratio applicable to any medical treatment. The actual amount administered, and the rate and time-course of administration, will depend on the nature and severity of the condition being treated. Prescription of treatment, e.g. decisions on dosage, timing, etc., is within the responsibility of general practitioners or specialists, and typically takes account of the disorder to be treated, the condition of the individual patient, the site of delivery, the method of administration and other factors known to practitioners. Examples of techniques and protocols can be found in *Remington's Pharmaceutical Sciences*.

[0076] Dosage may be adjusted appropriately to achieve desired drug levels, locally or systemically. Typically the active agents of the present invention exhibit their effect at a dosage range from about 0.001 mg/kg to about 250 mg/kg, preferably from about 0.01 mg/kg to about 100 mg/kg of the active ingredient, more preferably from about 0.05 mg/kg to about 75 mg/kg. A suitable dose can be administered in multiple sub-doses per day. Typically, a dose or sub-dose may contain from about 0.1 mg to about 500 mg of the active ingredient per unit dosage form. A more preferred dosage will contain from about 0.5 mg to about 100 mg of active ingredient per unit dosage form. Dosages are generally initiated at lower levels and increased until desired effects are achieved. In the event that the response in a subject is insufficient at such doses, even higher doses (or effective higher doses by a different, more localized delivery

route) may be employed to the extent that patient tolerance permits. Continuous dosing over, for example 24 hours or multiple doses per day are contemplated to achieve appropriate systemic levels of compounds.

[0077] For the treatment of pain, if the route of administration is directly to the CNS, the dosage contemplated is from about 1 ng to about 100 mg per day, preferably from about 100 ng to about 10 mg per day, more preferably from about 1  $\mu$ g to about 100  $\mu$ g per day. If administered peripherally, the dosage contemplated is somewhat higher, from about 100 ng to about 1000 mg per day, preferably from about 10  $\mu$ g to about 100 mg per day, more preferably from about 100  $\mu$ g to about 10 mg per day. If the conopeptide is delivered by continuous infusion (e.g., by pump delivery, biodegradable polymer delivery or cell-based delivery), then a lower dosage is contemplated than for bolus delivery.

[0078] Advantageously, the compositions are formulated as dosage units, each unit being adapted to supply a fixed dose of active ingredients. Tablets, coated tablets, capsules, ampoules and suppositories are examples of dosage forms according to the invention.

[0079] It is only necessary that the active ingredient constitute an effective amount, i.e., such that a suitable effective dosage will be consistent with the dosage form employed in single or multiple unit doses. The exact individual dosages, as well as daily dosages, are determined according to standard medical principles under the direction of a physician or veterinarian for use humans or animals.

[0080] The pharmaceutical compositions will generally contain from about 0.0001 to 99 wt. %, preferably about 0.001 to 50 wt. %, more preferably about 0.01 to 10 wt.% of the active ingredient by weight of the total composition. In addition to the active agent, the pharmaceutical compositions and medicaments can also contain other pharmaceutically active compounds. Examples of other pharmaceutically active compounds include, but are not limited to, analgesic agents, cytokines and therapeutic agents in all of the major areas of clinical medicine. When used with other pharmaceutically active compounds, the conopeptides of the present invention may be delivered in the form of drug cocktails. A cocktail is a mixture of any one of the compounds useful with this invention with another drug or agent. In this embodiment, a common administration vehicle (e.g., pill, tablet, implant, pump, injectable solution, etc.) would contain both the instant composition in combination supplementary potentiating agent. The individual drugs of the cocktail are each administered in therapeutically effective amounts. A therapeutically effective amount will be determined by the parameters described above; but, in

any event, is that amount which establishes a level of the drugs in the area of body where the drugs are required for a period of time which is effective in attaining the desired effects.

[0081] The practice of the present invention employs, unless otherwise indicated, conventional techniques of chemistry, molecular biology, microbiology, recombinant DNA, genetics, immunology, cell biology, cell culture and transgenic biology, which are within the skill of the art. See, e.g., Maniatis *et al.*, 1982; Sambrook *et al.*, 1989; Ausubel *et al.*, 1992; Glover, 1985; Anand, 1992; Guthrie and Fink, 1991; Harlow and Lane, 1988; Jakoby and Pastan, 1979; *Nucleic Acid Hybridization* (B. D. Hames & S. J. Higgins eds. 1984); *Transcription And Translation* (B. D. Hames & S. J. Higgins eds. 1984); *Culture Of Animal Cells* (R. I. Freshney, Alan R. Liss, Inc., 1987); *Immobilized Cells And Enzymes* (IRL Press, 1986); B. Perbal, *A Practical Guide To Molecular Cloning* (1984); the treatise, *Methods In Enzymology* (Academic Press, Inc., N.Y.); *Gene Transfer Vectors For Mammalian Cells* (J. H. Miller and M. P. Calos eds., 1987, Cold Spring Harbor Laboratory); *Methods In Enzymology*, Vols. 154 and 155 (Wu *et al.* eds.), *Immunochemical Methods In Cell And Molecular Biology* (Mayer and Walker, eds., Academic Press, London, 1987); *Handbook Of Experimental Immunology*, Volumes I-IV (D. M. Weir and C. C. Blackwell, eds., 1986); Riott, *Essential Immunology*, 6th Edition, Blackwell Scientific Publications, Oxford, 1988; Hogan *et al.*, *Manipulating the Mouse Embryo*, (Cold Spring Harbor Laboratory Press, Cold Spring Harbor, N.Y., 1986).

### EXAMPLES

[0082] The present invention is described by reference to the following Examples, which are offered by way of illustration and are not intended to limit the invention in any manner. Standard techniques well known in the art or the techniques specifically described below were utilized.

#### EXAMPLE 1

##### Isolation of $\mu$ -Conopeptides

[0083] Crude venom was extracted from venom ducts (Cruz *et al.*, 1976), and the components were purified as previously described (Cartier *et al.*, 1996). The crude extract from venom ducts was purified by reverse phase liquid chromatography (RPLC) using a Vydac C<sub>18</sub>

semi-preparative column (10 x 250 mm). Further purification of bioactive peaks was done on a Vydac C<sub>18</sub> analytical column (4.6 x 220 mm). The effluents were monitored at 220 nm. Peaks were collected, and aliquots were assayed for activity. Throughout purification, HPLC fractions were assayed by means of intracerebral ventricular (i.c.v.) injection into mice (Clark et al., 1981).

[0084] The amino acid sequence of the purified peptides were determined by standard methods. The purified peptides were reduced and alkylated prior to sequencing by automated Edman degradation on an Applied Biosystems 477A Protein Sequencer with a 120A Analyzer (DNA/Peptide Facility, University of Utah) (Martinez et al., 1995; Shon et al., 1994).

[0085] In accordance with this method, the  $\mu$ -conopeptides described as "isolated" in Table 1 were obtained. These  $\mu$ -conopeptides, as well as the other  $\mu$ -conopeptides and the  $\mu$ -conopeptide precursors set forth in Table 1 are synthesized as described in U.S. Patent No. 5,670,622.

## EXAMPLE 2

### Isolation of DNA Encoding $\mu$ -Conopeptides

[0086] DNA coding for  $\mu$ -conopeptides was isolated and cloned in accordance with conventional techniques using general procedures well known in the art, such as described in Olivera et al. (1996). Alternatively, cDNA libraries was prepared from *Conus* venom duct using conventional techniques. DNA from single clones was amplified by conventional techniques using primers which correspond approximately to the M13 universal priming site and the M13 reverse universal priming site. Clones having a size of approximately 300-500 nucleotides were sequenced and screened for similarity in sequence to known  $\mu$ -conotoxins. The DNA sequences and encoded propeptide sequences are set forth in Table 1. DNA sequences coding for the mature toxin can also be prepared on the basis of the DNA sequences set forth in Table 1. An alignment of the  $\mu$ -conopeptides of the present invention is set forth in Table 2.

TABLE 1

DNA and Amino Acid Sequences of  $\mu$ -Conopeptides and Precursors

Name:	Ar3.1
Species:	arenatus
Cloned:	Yes



**DNA Sequence:**

CAAGAAGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTCTTGACCATCTG  
TATGCTTCTGTTTCCCCTTACTGCTCTTCCGCTGGATGGGGATCAACCTGCAGACCG  
ACCTGCAGAGCGTATGCAGGACGACTTTATAACTGAGCATCATCCCCTGTTTGATCC  
5 TGTCAAACGGTGTTGCGAGAGGCCATGCAACATAGGATGCGTACCTTGTTGTTAATG  
ACCAGCTTTGTCATCGCGGCCTCATCAAGCGAATAAGTAAAACGATTGCAGT (SEQ  
ID NO:1)

**Translation:**

10 MMSKLGVFLTICMLLFPLTALPLDGDQPADRPAERMQDDFITEHHPLFDPVKRCCERPC  
NIGCVPC (SEQ ID NO:2)

**Toxin Sequence:**

15 Cys-Cys-Xaa1-Arg-Xaa3-Cys-Asn-Ile-Gly-Cys-Val-Xaa3-Cys-Cys-^ (SEQ ID NO:3)

**Name:** Ak3.1  
**Species:** atlanticus  
**Cloned:** Yes

20

**DNA Sequence:**

GGATCCATGATGTCTAAACTGGGAGTCTTGTTGACCATCTGTCTGCTTCTGTTTCCAC  
TTACTGCTCTTCCGCTGGATGAAGATCAACCGGTACACCGACCTGCAGAGCGTATGC  
AGGACATTTTCATCTGATCAACATCTCTTCTTTGATCTCATCAAACGGTGCTGCGAGT  
25 TGCCATGCGGGCCAGGCTTTTGCGTCCCTTGTTGCTGACATCAATAACGTGTTGATG  
ACCAACTTTCTCGAG (SEQ ID NO:4)

**Translation:**

30 GSMMSKLGVLITICLLLFPLTALPLDEDQPVHRPAERMQDISSDQHLFFDLIKRCCCLPC  
GPGFCVPCC (SEQ ID NO:5)

**Toxin Sequence:**

35 Cys-Cys-Xaa1-Leu-Xaa3-Cys-Gly-Xaa3-Gly-Phe-Cys-Val-Xaa3-Cys-Cys-^ (SEQ ID NO:6)

**Name:** A3.1  
**Species:** aurisiacus  
**Cloned:** Yes

**DNA Sequence:**

40 CAAGAGGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGTTGACCATCTG  
TTTGCTTCTGTTTCCCCTTACTGCTCTTCCGATGGATGGAGATCAATCTGTAGACCGA  
CCTGAAGAGCGTATGCAGGACGACATTTTCATCTGAGCAGCATCCCTTGTTTAATCAG  
AAAAGAATGTGTTGCGGCGAAGGCCGGAATGCCCCAGCTATTTTCAGAAACAGTCA  
45 GATTTGTCATTGTTGTTAAATGACAACGTGTCGATGACCAACTTCGTTATCACGACT  
AATGAATAAGTAAAACGATTGCAGT (SEQ ID NO:7)

**Translation:**

MMSKLGVLLTICLLLFPLTALPMDGDQSVDRPEERMQDDISSEQHPLFNQKRMCCGEG  
RKCPYSYFRNSQICHCC (SEQ ID NO:8)

5 **Toxin Sequence:**

Met-Cys-Cys-Gly-Xaa1-Gly-Arg-Lys-Cys-Xaa3-Ser-Xaa5-Phe-Arg-Asn-Ser-Gln-Ile-Cys-His-  
Cys-Cys-^ (SEQ ID NO:9)

10 **Name:** A3.2  
**Species:** aurisiacus  
**Cloned:** Yes

**DNA Sequence:**

15 GGATCCATGATGTCTAAACTGGGAGTCTTGTTGACCATCTGTTTGCTTCTGTTTCCCC  
TTACTGCTCTTCCGATCGATGGAGATCAATCTGTAGACCGACCTGCAGAGCGTATGC  
AGGATGACATTTTCATCTGAGCAGCATCGCTTGTTCAATCAGAAAAGAAGGTGCTGC  
CGGTGGCCATGCCCCCGACAAATCGACGGTGAATATTGTGGCTGTTGCCTTGGATGA  
TAACCGTGTTGATGACCAACTTTCTCGAG (SEQ ID NO:10)

20

**Translation:**

GSMMSKLGVLLTICLLLFPLTALPIDGDQSVDRPAERMQDDISSEQHRLFNQKRRCRW  
PCPRQIDGEYCGCCLG (SEQ ID NO:11)

25 **Toxin Sequence:**

Cys-Cys-Arg-Xaa4-Xaa3-Cys-Xaa3-Arg-Gln-Ile-Asp-Gly-Xaa1-Xaa5-Cys-Gly-Cys-Cys-Leu-#  
(SEQ ID NO:12)

30 **Name:** A3.3  
**Species:** aurisiacus  
**Cloned:** Yes

**DNA Sequence:**

35 GGATCCATGATGTCTAAACTGGGAGTCTTGTTGACCATCTGTCTACTTCTGTTTCCCC  
TTACTGCTTTTCCGATGGATGGAGATCAACCTGCAGACCAACCTGCAGATCGTATGC  
AGGACGACATTTTCATCTGAGCAGTATCCCTTGTTTGATAAGAGACAAAAGTGTTGCA  
CTGGGAAGAAGGGGTCTGCTCCGGCAAAGCATGCAAAAATCTCAAATGTTGCTCT  
GGACGATAACGTGTTGATGACCAACTTTCTCGAG (SEQ ID NO:13)

40

**Translation:**

GSMMSKLGVLLTICLLLFPLTAFPMDDGDQPADQPADRMQDDISSEQYPLFDKRQKCCT  
GKKGSCSGKACKNLKCCSGR (SEQ ID NO:14)

45 **Toxin Sequence:**

Xaa2-Lys-Cys-Cys-Thr-Gly-Lys-Lys-Gly-Ser-Cys-Ser-Gly-Lys-Ala-Cys-Lys-Asn-Leu-Lys-  
Cys-Cys-Ser-# (SEQ ID NO:15)

**Name:** A3.4  
**Species:** aurisiacus  
**Cloned:** Yes

5

**DNA Sequence:**

GGATCCATGATGTCTAAACTGGGAGTCTTGCTGACCATCTGTCTGCTTCTGTTTCCAC  
TTACTGCTGTTCCGCTGGATGGAGATCAACCTCTAGACCGACACGCGGAGCGTATGC  
10 ATGATGGCATTTACCTAAACGCCATCCCTGGTTTGATCCCGTCAAACGGTGTGCA  
AGGTGCAATGCGAGTCTTGACCCCTTGTTGCTAACGTGTTGATGACCAACTTTCTC  
GAG (SEQ ID NO:16)

**Translation:**

15 GSMMSKLGVLLTICLLFPLTAVPLDGDQPLDRHAERMHDGISPKRHPWFDPVKRCK  
VQCESCTPCC (SEQ ID NO:17)

**Toxin Sequence:**

20 Cys-Cys-Lys-Val-Gln-Cys-Xaa1-Ser-Cys-Thr-Xaa3-Cys-Cys-^ (SEQ ID NO:18)

**Name:** Bn3.1  
**Species:** bandanus  
**Cloned:** Yes

25

**DNA Sequence:**

GGATCCATGATGTCTAAACTGGGAGTCTTGTTGACCATCTGTATGCTTCTGTTTCCCC  
TCACTGCTCTTCCGATGGATGGAGATCAACCTGCAGACCGACCTGCAGAGCGTAGT  
CAGGACGTTTCATCTGAACAGCATCCCTTGTTTGATCCCGTCAAACGGTGTGCAAC  
30 TGGCCATGCTCCATGGGATGCATCCCTTGTTGCTACTATTAATAACGTGTTGATGAC  
CAACTTTCTCGAG (SEQ ID NO:19)

**Translation:**

35 GSMMSKLGVLLTICMLLFPLTALPMDGDQPADRPAERSQDVSSEQHPLFDPVKRCCNW  
PCSMGCIPCCYY (SEQ ID NO:20)

**Toxin Sequence:**

40 Cys-Cys-Asn-Xaa4-Xaa3-Cys-Ser-Met-Gly-Cys-Ile-Xaa3-Cys-Cys-Xaa5-Xaa5-^ (SEQ ID  
NO:21)

**Name:** Bt3.1  
**Species:** betulinus  
**Cloned:** Yes

45

**DNA Sequence:**

CAAGAGGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGTTGACCTTCTG

TCTGCTTCTGTTTCCCCTGACTGCTCTTCCGCTGGATGAAGATCAACCTGCAGACCG  
ACCTGCAGAGCGTATGCAGGACATTTTCATCTGAACAGCATCCCTTGTTTGATCCCGT  
CAAACGGTGTTGCGAATTGCCATGCCATGGATGCGTCCCTTGTTGCTGGCCTTAATA  
ACGTGTGGATGACCAACTGTGTTATCACGGCCACGTCAAGTGTCTAATGAATAAGT  
5 AAAATGATTGCAGT (SEQ ID NO:22)

**Translation:**

MMSKLGVLLTFCLLLFPLTALPLDEDQPADRPAERMQDISSEQHPLFDPVKRCCELPC  
10 GCVPCWP (SEQ ID NO:23)

**Toxin Sequence:**

Cys-Cys-Xaa1-Leu-Xaa3-Cys-His-Gly-Cys-Val-Xaa3-Cys-Cys-Xaa4-Xaa3-^ (SEQ ID NO:24)

15 **Name:** Bt3.2  
**Species:** betulinus  
**Cloned:** Yes

**DNA Sequence:**

20 CAAGAGGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGTTGACCTTCTG  
TCTGCTTCTGTTTCCCCTGACTGCTCTTCCGCTGGATGAAGATCAACCTGCAGACCG  
ACATGCAGAGCGTATGCAGGACATTTACCTGAACAGCATCCCTCGTTTGATCCCGT  
CAAACGGTGTTGCGGGCTGCCATGCAATGGATGCGTCCCTTGTTGCTGGCCTTCATA  
ACGTGTGGACGACCAACTTTGTTATCACGGCCACGTCAAGTGTCTGATGAATAAGTA  
25 AAACGATTGCAGT (SEQ ID NO:25)

**Translation:**

MMSKLGVLLTFCLLLFPLTALPLDEDQPADRHARMQDISPEQHPSFDPVKRCCGLPCN  
30 GCVPCWP (SEQ ID NO:26)

**Toxin Sequence:**

Cys-Cys-Gly-Leu-Xaa3-Cys-Asn-Gly-Cys-Val-Xaa3-Cys-Cys-Xaa4-Xaa3-Ser-^ (SEQ ID  
NO:27)

35 -----

**Name:** Bt3.3  
**Species:** betulinus  
**Cloned:** Yes

40

**DNA Sequence:**

CAAGAGGGATCGATAGCAGTTCATGATGTTTAAACTGGGAGTCTTGTTGACCATCTA  
TATGCTTCTGTTTCCCTTTACTGCTCTTCCGCTGGATGGAGATCAACCTGCAGACCAA  
CCTCTAGAGCGCATGCAGTATGACATGTTACGTGCAGTGAATCCCTGGTTTGATCCC  
45 GTCAAAAGGTGCTGCTCGAGGAACTGCGCAGTATGCATCCCTTGTTGCCCGAATTGG  
CCAGCTTGATTATCGCGGCCAAGAGTCTAATGAATAAGTAAAACGATTGCAGT (SEQ  
ID NO:28)

**Translation:**

MMFKLGVLLTIYMLLPFTALPLDGDQPADQPLERMQYDMLRAVNPWFDPVKRCCSR  
NCAVCIPCCPNWPA (SEQ ID NO:29)

5

**Toxin Sequence:**

Cys-Cys-Ser-Arg-Asn-Cys-Ala-Val-Cys-Ile-Xaa3-Cys-Cys-Xaa3-Asn-Xaa4-Xaa3-Ala-^ (SEQ  
ID NO:30)

10

**Name:** Bu3.1  
**Species:** bullatus  
**Cloned:** Yes

15 **DNA Sequence:**

CAAGAAGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGTTGACCATCTG  
TCTGCTTCTGTTTCCCCCTTTTGCTCTTCCGCAGGATGGAGATCAACCTGCAGACCGA  
CCTGCAGAGCGTATGCAGGACGACATTTTCATCTGAGCAGAATTCCTTGCTTGAGAA  
GAGAGTTACTGACAGGTGCTGCAAAGGGAAGAGGGAATGCGGCAGATGGTGCAGA  
20 GATCACTCGCGTTGTTGCGGTCGACGATAAGCTGTTGATGACCAGCTTTGTTATCAC  
GGCTACATCAAGTGTCTAGTGAATAAGTAAAATGATTGCAGT (SEQ ID NO:31)

**Translation:**

MMSKLGVLLTICLLLFPLFALPQDGDQPADRPAERMQDDISSEQNSLLEKRVTDRCCKG  
25 KRECGRWCRDHSRCCGRR (SEQ ID NO:32)

**Toxin Sequence:**

Val-Thr-Asp-Arg-Cys-Cys-Lys-Gly-Lys-Arg-Xaa1-Cys-Gly-Arg-Xaa4-Cys-Arg-Asp-His-Ser-  
Arg-Cys-Cys-# (SEQ ID NO:33)

30

**Name:** Bu3.1A  
**Species:** bullatus  
**Cloned:** Yes

35

**DNA Sequence:**

CAAGAGGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGTTGACCATCTG  
TCTGCTTCTGTTTCCCCCTTTTGCTCTTCCGCAGGATGGAGATCAACCTGCAGACCGA  
CCTGCAGAGCGTATGCAGGATGACATTTTCATCTGAGCAGAATCCCTTGCTTGAGAA  
40 GAGAGTTGGTGACAGGTGCTGCAAAGGGAAGAGGGGGTGCGGCAGATGGTGCAGA  
GATCACTCACGTTGTTGCGGTCGACGATAACGTGTTGATGACCAGCTTTGTTATCAC  
GGCTACATCAAGTGTCTTAGTGATTAAGTAAAACGATTGCAGT (SEQ ID NO:34)

**Translation:**

45 MMSKLGVLLTICLLLFPLFALRQDGDQPADRPAERMQDDISSEQNPLLEKRVGDRCK  
GKRGCGRWCRDHSRCCGRR (SEQ ID NO:35)

**Toxin Sequence:**

Val-Gly-Asp-Arg-Cys-Cys-Lys-Gly-Lys-Arg-Gly-Cys-Gly-Arg-Xaa4-Cys-Arg-Asp-His-Ser-Arg-Cys-Cys-# (SEQ ID NO:36)

5

**Name:** Bu3.2  
**Species:** bullatus  
**Cloned:** Yes

10 **DNA Sequence:**

CAAGAAGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGTTGACCATCTG  
TCTGCTTCTGTTTCCCTTTTGTCTTCCGCAGGATGGAGATCAACCTGCAGACCGA  
CCTGCAGAGCGTATGCAGGACGACATTTTCATCTGAGCAGAATCCCTTGCTTGAGAA  
GAGAGTTGGTGAAAGGTGCTGCAAAAACGGGAAGAGGGGGTGCAGGAGATGGTGC  
15 AGAGATCACTCACGTTGTTGCGGTCGACGATAACGTGTTGATGACCGAGGCTTTCGT  
TATCACGGCTACATCAAGTGTCTAGTGAATAAGTAAAACGATTGCAGT (SEQ ID  
NO:37)

**Translation:**

20 MMSKLGVLTTICLLLFPLFALPDGDQPADRPAERMQDDISSEQNPLLEKRVGERCCKN  
GKRGCSRWCRDHSRCCGRR (SEQ ID NO:38)

**Toxin Sequence:**

25 Val-Gly-Xaa1-Arg-Cys-Cys-Lys-Asn-Gly-Lys-Arg-Gly-Cys-Gly-Arg-Xaa4-Cys-Arg-Asp-His-  
Ser-Arg-Cys-Cys-# (SEQ ID NO:39)

**Name:** Bu3.3  
**Species:** bullatus  
30 **Cloned:** Yes

**DNA Sequence:**

CAAGAGGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGTTGACCATCTG  
TCTGCTTCTGTTTCCCTTTTGTCTTCCGCAGGACGGAGATCAACCTGCAGACCG  
35 ACCTGCAGAGCGTATGCAGGACGACCTTTCATCTGAGCAGCATCCCTTGTTTGAGAA  
GAGAATTGTTGACAGGTGCTGCAACAAAGGGAACGGGAAGAGGGGGTGCAGCAGA  
TGGTGCAGAGATCACTCACGTTGTTGCGGTCGACGATGAACTGTTGATGACCGAGG  
CTTTGGTTATCACGGCTACATCAAGTGTCTAGTGAATAAGTAAAACGATTGCAGT  
(SEQ ID NO:40)

40

**Translation:**

MMSKLGVLTTICLLLFPLFALPDGDQPADRPAERMQDDLSEQHPLFEKRIVDRCCNK  
GNGKRGCSRWCRDHSRCCGRR (SEQ ID NO:41)

45 **Toxin Sequence:**

Ile-Val-Asp-Arg-Cys-Cys-Asn-Lys-Gly-Asn-Gly-Lys-Arg-Gly-Cys-Ser-Arg-Xaa4-Cys-Arg-Asp-His-Ser-Arg-Cys-Cys-# (SEQ ID NO:42)

**Name:** Bu3.4  
**Species:** bullatus  
**Cloned:** Yes

**DNA Sequence:**

CAAGAAGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGTTGACCATCTG  
TCTGCTTCTGTTTCCCCTTTTTGCTCTTCCGCAGGATGGAGATCAACCTGCAGACCGA  
10 CCTGCTGAGCGTATGCAGGACGACATTTTCATCTGAGCGGAATCCCTTGTTTGAGAAG  
AGCGTTGGTTTATATTGCTGCCGACCCAAACCCAACGGGCAGATGATGTGCGACAG  
ATGGTGCGAAAAAACTCACGTTGTTGCGGTCGACGATAATGTGTTGATGACCAGC  
TTTGTTATCAAGGCTACATCAAGTATCTAGTGAATAAGTAAACGATTGCAGT (SEQ  
ID NO:43)

**Translation:**

MMSKLGVLLTICLLLFPLFALPDGDQPADRPAERMQDDISSERNPLFEKSVGLYCCRP  
KPNGQMMCDRWCEKNSRCCGRR (SEQ ID NO:44)

**Toxin Sequence:**

Val-Gly-Leu-Xaa5-Cys-Cys-Arg-Xaa3-Lys-Xaa3-Asn-Gly-Gln-Met-Met-Cys-Asp-Arg-Xaa4-  
Cys-Xaa1-Lys-Asn-Ser-Arg-Cys-Cys-# (SEQ ID NO:45)

-----  
**Name:** Bu3.5  
**Species:** bullatus  
**Cloned:** Yes

**DNA Sequence:**

CAAGAAGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTTTTGTTGACCATCTG  
TCTGCTTCTGTTTCCCCTTACTGCTCTTCCGATGGATGGAGATCAATCTGTAGACCGA  
CCTGCAGAACGTATGCAGGACGACCTTTCATCTGAGCAGCATCCCTTGTTTGTTTCAG  
AAAAGAAGGTGTTGCGGCGAAGGCTTGACATGCCCCAGATATTGGAAAAACAGTCA  
35 GATTTGTGCTTGTTGTTAAATGACAACGTGTGCGATGACCAACTTCGGTATCACGACT  
ACGCCAAGTGTCTAATGAATAAGTAAACGATTGCAGT (SEQ ID NO:46)

**Translation:**

MMSKLGVLLTICLLLFPLTALPMDGDQSVDRPAERMQDDLSEQHPLFVQKRRCCGEG  
40 LTCPRYWKNSQICACC (SEQ ID NO:47)

**Toxin Sequence:**

Arg-Cys-Cys-Gly-Xaa1-Gly-Leu-Thr-Cys-Xaa3-Arg-Xaa5-Xaa4-Lys-Asn-Ser-Gln-Ile-Cys-Ala-  
Cys-Cys-^ (SEQ ID NO:48)

**Name:** Bu3.5A  
**Species:** bullatus  
**Cloned:** Yes

5 **DNA Sequence:**

CAAGAGGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGTTGACCATCTG  
TCTGCTTCTGTTTCCCCTTTTGTCTTCCGCAGGATGGAGATCAACCTGCAGACCGA  
CCTGCTGAGCGTATGCAGGACGACATTTTCATCTGAGCAGGATCCCTTGTTTGTTGAG  
AAAAGAAGGTGTTGCGGCGAAGGCTTGACATGCCCCAGATATTGGAAAAACAGTCA  
10 GATTTGTGCTTGTTGTTAAATGACAACGTGTGATGACCAACTTCGGTATCACGACTA  
CGCCAAGTGTCTAATGAATAAGTAAAACGATTGCAGT (SEQ ID NO:49)

**Translation:**

MMSKLGVLITICLLLFPLFALPQDGDQPADRPAERMQDDISSEQDPLFVQKRRCCGEGL  
15 TCPRYWKNSQICACC (SEQ ID NO:50)

**Toxin Sequence:**

Arg-Cys-Cys-Gly-Xaa1-Gly-Leu-Thr-Cys-Xaa3-Arg-Xaa5-Xaa4-Lys-Asn-Ser-Gln-Ile-Cys-Ala-  
Cys-Cys-^ (SEQ ID NO:51)

20

**Name:** Cp3.1  
**Species:** capitaneus  
**Cloned:** Yes

25

**DNA Sequence:**

GGATCCATGATGTCTAAACTGGGAGTCTTGGTGACCATCTGCCTGCTTCTGTTTCCC  
CTTGCTGCTTTTCCACTGGATGGAAATCAACCTGCAGACCACCCTGCAAAGCGTACG  
CAAGATGACAGTTCAGCTGCCCTGATCAATACCTGGATTGATCATTCCCATTCTTGC  
30 TGCAGGGACTGCGGTGAAGATTGTGTTGGTTGTTGCCGGTAACGTGTTGATGACCAA  
CTTTCTCGAG (SEQ ID NO:52)

**Translation:**

GSMMSKLGVLVTICLLLFPLAFLDGNQPADHPAKRTQDDSSAALINTWIDHSHSCCR  
35 DCGEDCVGCCR (SEQ ID NO:53)

**Toxin Sequence:**

Ser-Cys-Cys-Arg-Asp-Cys-Gly-Xaa1-Asp-Cys-Val-Gly-Cys-Cys-Arg-^ (SEQ ID NO:54)

40

**Name:** Ca3.1  
**Species:** characteristicus  
**Cloned:** Yes

45 **DNA Sequence:**

CAAGAGGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGTTGACCATCTG  
TCTGCTTCTGTTTCCCCTTACTGCTCTTCCAATGGATGGAGATCAACCTGCAGACCA



ACCTGCAGATCGTATGCAGGACGACATTTTCATCTGAGCAGTATCCCTTGTTTGATAT  
GAGAAAAAGGTGTTGCGGCCCCGGCGGTTTCATGCCCCGTATATTTTCAGAGACAATT  
TTATTTGTGGTTGTTGTTAAATGACAACGTGTCGATGACCAACTTCATTATCACGAC  
TACGCCAAGTGTCTAATGAATAAGTAAAATGATTGCAGT (SEQ ID NO:55)

5

**Translation:**

MMSKLGVLITICLLLFPLTALPMDGDQPADQPADRMQDDISSEQYPLFDMRKRC CGPG  
GSCP VYFRDNFICGCC (SEQ ID NO:56)

10

**Toxin Sequence:**

Cys-Cys-Gly-Xaa3-Gly-Gly-Ser-Cys-Xaa3-Val-Xaa5-Phe-Arg-Asp-Asn-Phe-Ile-Cys-Gly-Cys-  
Cys-^ (SEQ ID NO:57)

15

**Name:** Ca3.2  
**Species:** characteristic  
**Cloned:** Yes

**DNA Sequence:**

20

CAAGAGGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGTTGACCATCTG  
TCTGCTTCTGTTTCCCCTTACTGCTCTTCCGATGGATGGAGATGAACCTGCAAACCG  
ACCTGTGCGAGCGTATGCAGGACAACATTTTCATCTGAGCAGTATCCCTTGTTTGAGAA  
GAGACGAGATTGTTGCACTCCGCCGAAGAAATGCAAAGACCGACAATGCAAACCCC  
AGAGATGTTGCGCTGGACGATAACGTGTTGATGACCAACTTTATCACGGCTACGTCA  
25 AGTGTTTAGTGAATAAGTAAAATGATTGCAGT (SEQ ID NO:58)

**Translation:**

MMSKLGVLITICLLLFPLTALPMDGDEPANRPVERMQDNISSEQYPLFEKRRDCCTPPK  
KCKDRQCKPQRCCAGR (SEQ ID NO:59)

30

**Toxin Sequence:**

Arg-Asp-Cys-Cys-Thr-Xaa3-Xaa3-Lys-Lys-Cys-Lys-Asp-Arg-Gln-Cys-Lys-Xaa3-Gln-Arg-  
Cys-Cys-Ala-# (SEQ ID NO:60)

35

**Name:** Ca3.3  
**Species:** characteristic  
**Cloned:** Yes

40

**DNA Sequence:**

CAAGAGGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGTTGACCATCTG  
TCTGCTTCTGTTTCCCCTTACTGCTCTTCCACTGGATGGAGATCAACCTGCAGATCAA  
TCTGCAGAGCGACCTGCAGAGCGTACGCAGGACGACATTCAGCAGCATCCGTTATA  
TGATCCGAAAAGAAGGTGTTGCCGTTATCCATGCCCCGACAGCTGCCACGGATCTTG  
45 CTGCTATAAGTGATAACATGTTGATGGCCAGCTTTGTTATCACGGCCACGTCAAGTG  
TCTTAATGAATAAGTAAAACGATTGCAGT (SEQ ID NO:61)

**Translation:**

MMSKLGVLLTICLLLFPLTALPLDGDQPADQSAERPAERTQDDIQHPLYDPKRRCCRY  
PCPDSCHGSCCYK (SEQ ID NO:62)

5 **Toxin Sequence:**

Arg-Cys-Cys-Arg-Xaa5-Xaa3-Cys-Xaa3-Asp-Ser-Cys-His-Gly-Ser-Cys-Cys-Xaa5-Lys-^ (SEQ  
ID NO:63)

10 **Name:** Ca3.4  
**Species:** characteristic  
**Cloned:** Yes

**DNA Sequence:**

15 CAAGAGGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGCCTTGTTGACCATCT  
GTCTACTTCTGTTTTCCCTTACTGCTGTTCCGCTGGATGGAGATCAACATGCAGACC  
AACCTGCACAGCGTCTGCAGGACCGCATTCCAACCTGAAGATCATCCCTTATTTGATC  
CCAACAAACGGTGTTGCCCGCCGGTGGCATGCAACATGGGATGCAAGCCTTGTTGT  
20 GGATGACCAGCTTTGTTATCGCGGTCTTCATGAAGTGTCTTAATGAATAAGTAAAAT  
GATTGCAGT (SEQ ID NO:64)

**Translation:**

MMSKLGALLTICLLLFSLTAVPLDGDQHADQPAQRLQDRIPTEDHPLFDPNKRCCPPVA  
CNMGCKPCCG (SEQ ID NO:65)

25

**Toxin Sequence:**

Cys-Cys-Xaa3-Xaa3-Val-Ala-Cys-Asn-Met-Gly-Cys-Lys-Xaa3-Cys-Cys-# (SEQ ID NO:66)

30 **Name:** Ca3.5  
**Species:** characteristic  
**Cloned:** Yes

**DNA Sequence:**

35 CAAGAGGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGCCTTGTTGACCATCT  
GTCTACTTCTGTTTTCCCTAACTGCTGTTCCGCTGGATGGAGATCAACATGCAGACC  
AACCTGCAGAGCGTCTGCATGACCGCCTTCCAACCTGAAAATCATCCCTTATATGATC  
CCGTCAAACGGTGTTGCGATGATTCGGAATGCGACTATTCTTGCTGGCCTTGCTGTA  
TGTTTGGATAACCTTTGTTATCGCGGCCTCATCAAGTGTCTAATGAATAAGTAAAAC  
40 GATTGCAGT (SEQ ID NO:67)

**Translation:**

MMSKLGALLTICLLLFSLTAVPLDGDQHADQPAERLHDRLPTENHPLYDPVKRCCDDSE  
CDYSCWPCCMFG (SEQ ID NO:68)

45

**Toxin Sequence:**

Cys-Cys-Asp-Asp-Ser-Xaa1-Cys-Asp-Xaa5-Ser-Cys-Xaa4-Xaa3-Cys-Cys-Met-Phe-# (SEQ ID

NO:69)

5     **Name:**       Ca3.6  
      **Species:**   characteristicus  
      **Cloned:**    Yes

**DNA Sequence:**

10   GGATCCATGATGTCTAAACTGGGAGTCTTGTTGACCATCTGTCTGCTTCTGTTTCCCC  
     TTACTGCTGTTCCGCTGGATGGAGATCAACCTGCAGACCGACCTGCAGAGCGTAAG  
     CAGGACGTTTCATCTGAACAGCATCCCTTCTTTGATCCCGTCAAACGGTGTGCCGC  
     CGGTGTTACATGGGATGCATCCCTTGTTGCTTTTAAACGTGTTGATGACCAACTTTCTC  
     GAG (SEQ ID NO:70)

15   **Translation:**

     GSMMSKLGVLITICLLLFPLTAVPLDGDQPADRPAERKQDVSSEQHPFFDPVKRCCRRRC  
     YMGCI PCCF (SEQ ID NO:71)

**Toxin Sequence:**

20   Cys-Cys-Arg-Arg-Cys-Xaa5-Met-Gly-Cys-Ile-Xaa3-Cys-Cys-Phe-^ (SEQ ID NO:72)

**Name:**       Cr3.1  
     **Species:**   circumciscus  
25   **Cloned:**    Yes

**DNA Sequence:**

30   CAAGAAGGATCGATAGCAGTTCATGATGTCTAAACTGGGGGTATTGTTGACCATCT  
     GTCTGCTTCTGTTTCCCCTTACTGCTCTTCCAATGGATGGAGATCAACCTGCAGACC  
     AACCTGCAGATCGTATGCAGGACGACATTTTCATCTGAGCAGTATCCCTTGTTTGATA  
     AGAGACGAAAGTGTTGCGGCAAAGACGGGCCATGCCCCAAATATTTCAAAGACAAT  
     TTTATTTGTGGTTGTTGTTAAATGACAACGTGTCGATGACCAACTTCGTTATCACGAT  
     TCGCCAAGTGTCTTAATGAATAAGTAAAATGATTGCAGT (SEQ ID NO:73)

35   **Translation:**

     MMSKLGVLITICLLLFPLTALPMDGDQPADQPADRMQDDISSEQYPLFDKRRKCCGKD  
     GPCPKYFKDNFICGCC (SEQ ID NO:74)

**Toxin Sequence:**

40   Arg-Lys-Cys-Cys-Gly-Lys-Asp-Gly-Xaa3-Cys-Xaa3-Lys-Xaa5-Phe-Lys-Asp-Asn-Phe-Ile-Cys-  
     Gly-Cys-Cys-^ (SEQ ID NO:75)

45   **Name:**       Da3.1  
     **Species:**   dalli  
     **Cloned:**    Yes

**DNA Sequence:**

CAAGAGGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGCCTTGTTGACCATCT  
GTCTACTTCTGTTTTCCCTAACTGCTGTTCCGCTGGATGGAGATCAACATGCAGACC  
AACCTGCAGAGCGTCTGCAGGACCGCCTTCCAACCTGAAAATCATCCCTTATATGATC  
5 CCGTCAAACGGTGTTGCGATGATTCGGAATGCGACTATTCTTGCTGGCCTTGCTGTA  
TTTTATCATAACCTTTGTTATCGCGGCCTCATCAAGTGTCAAATGAATAAGTAAAAT  
GATTGCAGT (SEQ ID NO:76)

**Translation:**

10 MMSKLGALLTICLLLFSLTAVPLDGDQHADQPAERLQDRLPTENHPLYDPVKRCCDDSE  
CDYSCWPCCILS (SEQ ID NO:77)

**Toxin Sequence:**

15 Cys-Cys-Asp-Asp-Ser-Xaa1-Cys-Asp-Xaa5-Ser-Cys-Xaa4-Xaa3-Cys-Cys-Ile-Leu-Ser-^ (SEQ  
ID NO:78)

Name: Da3.2

Species: dalli

20 Cloned: Yes

**DNA Sequence:**

CAAGAGGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGTTGACCATTG  
TCTACTTCTGTTTCCCTTACTGCTGTTCCACTGGATGGAGATCAGCCTGCAGACCG  
25 ACCTGCAGAGCGTATGCAGGACGGCATTTCATCTGAACATCATCCATTTTTTGGATTC  
CGTCAAAAAGAAACAACAGTGTTGCCCCGCGGTGGCATGCAACATGGGATGCGAGC  
CTTGTTGTGGATGACCAGCTTTGTTATCGCGGCTCATGAAGTGCCTAATGAATAAG  
TAAAACGATTGCAGT (SEQ ID NO:79)

**Translation:**

30 MMSKLGVLLTICLLLFPLTAVPLDGDQPADRPAERMQDGISSEHHPFFDSVKKKQQCCP  
PVACNMGCEPCCG (SEQ ID NO:80)

**Toxin Sequence:**

35 Xaa2-Gln-Cys-Cys-Xaa3-Xaa3-Val-Ala-Cys-Asn-Met-Gly-Cys-Xaa1-Xaa3-Cys-Cys-# (SEQ  
ID NO:81)

Name: Da3.3

40 Species: dalli

Cloned: Yes

**DNA Sequence:**

45 CAAGAAGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGTTGATCATATG  
TCTATTTCTGTTTCCCTTACTGCTGTTCACTCAATGGAGATCAGCCTGCAGACCAA  
TCTGCAGAGCGTATGCAGGACAAAATTCATCTGAACATCATCCCTTTTTTGGATCCC  
GTCAAACGTTGTTGCAACGCGGGTTTTGCCGCTTCGGATGCACGCCTTGTTGTTGG

TGACCAGCTTTGTTATCGCGGCCTCATCAAGTGTCTAATGAATAAGTAAAATGATTG  
CAGT (SEQ ID NO:82)

**Translation:**

5 MMSKLGVLIIICLFLFPLTAVQLNGDQPADQSAERMQDKISSEHHPFFDPVKRCCNAGF  
CRFGCTPCCW (SEQ ID NO:83)

**Toxin Sequence:**

10 Cys-Cys-Asn-Ala-Gly-Phe-Cys-Arg-Phe-Gly-Cys-Thr-Xaa3-Cys-Cys-Xaa4-^ (SEQ ID NO:84)

Name: Di3.1  
Species: distans  
Cloned: Yes

15

**DNA Sequence:**

CAAGAGGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGCTGACCATCTT  
TCTGCTTCTGTTTCCCTTACTGCTGTTCCGCTGGATGGAGATCAACCCGCAGACGG  
ACTTGCAGAGCGCATGCAGGACGACAGTTCAGCTGCACTGATTAGAGACTGGCTTC  
20 TTCAAACCCGACAGTGTGTGTGCATCCATGCCCATGCACGCCTTGCTGTAGATGAC  
CAGCTTTGTCATCGCGGCTACGTCAAGTATCTAATGAATAAGTAAAGTAAACGATTG  
CAGT (SEQ ID NO:85)

**Translation:**

25 MMSKLGVLITIFLLFPLTAVPLDGDQPADGLAERMQDDSSAALIRDWLLQTRQCCVH  
PCPCTPCCR (SEQ ID NO:86)

**Toxin Sequence:**

30 Xaa2-Cys-Cys-Val-His-Xaa3-Cys-Xaa3-Cys-Thr-Xaa3-Cys-Cys-Arg-^ (SEQ ID NO:87)

Name: E3.1  
Species: ermineus  
Cloned: Yes

35

**DNA Sequence:**

ACCTCAAGAGGGATCGATCGCAGTTCATGATGTCTAAACTGGGAGCCTTGTTGACC  
ATCTGTCTGCTTCTGTTTCCATTACTGCTCTTCTGATGGATGGAGATCAGCCTGCAG  
ACCGACCTGCAGAGCGTACGGAGGATGACATTTTCATCTGACTACATTCCTTGTGCA  
40 GTTGGCCATGCCCCGATACTCCAACGGTAAACTTGTTTGTGTTTGTGTCCTTGGATG  
ATAATGTGTTGATGACCAACTTTGTTATCACGGCTACGTCAAGTGTCTACTGAATAA  
GTAAATGATTGCAGTA (SEQ ID NO:88)

**Translation:**

45 MMSKLGALLTICLLLPITALLMDGDQPADRPAERTEDDISSDYIPCCSWPCPRYSNGKL  
VCFCLG (SEQ ID NO:89)

**Toxin Sequence:**

Cys-Cys-Ser-Xaa4-Xaa3-Cys-Xaa3-Arg-Xaa5-Ser-Asn-Gly-Lys-Leu-Val-Cys-Phe-Cys-Cys-Leu-# (SEQ ID NO:90)

5

**Name:** Ge3.2  
**Species:** generalis  
**Cloned:** Yes

10

**DNA Sequence:**

GGATCCATGATGTCTAAACTGGGAGTCTTGTTGACCATCTGTCTGGTTCTGTTTCCCC  
TTACTGCTCTTCCACTGGATGGAGAACAACCTGTAGACCGACATGCCGAGCATATGC  
AGGATGACAATTCAGCTGCACAGAACCCCTGGGTTATTGCCATCAGACAGTGTTC  
ACGTTCTGCAACTTTGGATGCCAACCTTGTTGCCTCACCTGATAACGTGTTGATGAC  
CAACTTTCTCGAG (SEQ ID NO:91)

15

**Translation:**

GSMMSKLGVLITICLVLFPLTALPLDGEQPVDRHAEHMQDDNSAAQNPWVIAIRQCCT  
FCNFGCQPCCLT (SEQ ID NO:92)

20

**Toxin Sequence:**

Xaa2-Cys-Cys-Thr-Phe-Cys-Asn-Phe-Gly-Cys-Gln-Xaa3-Cys-Cys-Leu-Thr-^ (SEQ ID NO:93)

25

**Name:** Ge3.3  
**Species:** generalis  
**Cloned:** Yes

**DNA Sequence:**

30

GGATCCATGATGTCTAAACTGGGAGTCTTGTTGACCATCTGTCTGGTTCTGTTTCCCC  
TTACTGCTCTTCCACTGGATGGAGAACAACCTGTAGACCGACATGCCGAGCATATGC  
AGGATGACAATTCAGCTGCACAGAACCCCTGGGTTATTGCCATCAGACAGTGTTC  
ACGTTCTGCAACTTTGGATGCCAGCCTTGTTGCGTCCCCTGATAACGTGTTGATGAC  
CAACTTTCTCGAG (SEQ ID NO:94)

35

**Translation:**

GSMMSKLGVLITICLVLFPLTALPLDGEQPVDRHAEHMQDDNSAAQNPWVIAIRQCCT  
FCNFGCQPCCV (SEQ ID NO:95)

40

**Toxin Sequence:**

Xaa2-Cys-Cys-Thr-Phe-Cys-Asn-Phe-Gly-Cys-Gln-Xaa3-Cys-Cys-Val-Xaa3-^ (SEQ ID NO:96)

45

**Name:**  $\mu$ -GIIIA  
**Species:** geographus  
**Cloned:** Yes

**DNA Sequence:**

5 GTCGACTCTAGAGGATCCGACAACAAAGAGTCAACCCCACTGCCACGTCAAGAGCG  
AAGCGCCACAGCTAAGACAAGAGGGATCGATAGCAGTTCATGATGTCTAAACTGGG  
AGTCTTGTGACCATCTGTCTGCTTCTGTTTCCCCTTACTGCTCTTCCGATGGATGGA  
10 GATGAACCTGCAAACCGACCTGTCGAGCGTATGCAGGACAACATTTTCATCTGAGCA  
GTATCCCTTGTTTGAGAAGAGACGAGATTGTTGCACTCCGCCGAAGAAATGCAAAG  
ACCGACAATGCAAACCCAGAGATGTTGCGCTGGACGATAACGTGTTGATGACCAA  
CTTTATCACGGCTACGTCAAGTGTTTAGTGAATAAGTAAAATGATTGCAGTCTTGCT  
CAGATTTGCTTTTGTGTTTTGGTCTAAAGATCAATGACCAAACCGTTGTTTTGATGCG  
15 GATTGTCATATATTTCTCGATTCCAATCCAACACTAGATGATTTAATCACGATAGAT  
TAATTTTCTATCAATGCCTTGATTTTTCGTCTGTCATATCAGTTTTGTTTATATTTATT  
TTTTCGTCACTGTCTACACAAACGCATGCATGCACGCATGCACGCACACACGCACGC  
ACGCTCGCACAAACATGCGCGCGCACGCACACACACACACACACACAAACACA  
20 CACACAAGCAATCACACAATTATTGACATTATTTATTTATTCATTGATGTATTTGTTA  
TTCGTTTGCTTGTTTTTAGAATAGTTTGAGGCCGTCTTTTTGGATTTATTTGAACTGC  
TTTATTGTATACGAGTACTTCGTGCTTTGAAACACTGCTGAAAATAAAACAAACACT  
GACGTAGC (SEQ ID NO:97)

**Translation:**

20 MMSKLGVLITICLLLFPLTALPMDGDEPANRPVERMQDNISSEQYPLFEKRRDCCTPPK  
KCKDRQCKPQRCCAGR (SEQ ID NO:98)

**Toxin Sequence:**

25 Arg-Asp-Cys-Cys-Thr-Xaa3-Xaa3-Lys-Lys-Cys-Lys-Asp-Arg-Gln-Cys-Lys-Xaa3-Gln-Arg-  
Cys-Cys-Ala-# (SEQ ID NO:99)

30 **Name:**  $\mu$ -GIIB  
**Species:** geographus  
**Isolated:** Yes  
**Cloned:** Yes

**DNA Sequence:**

35 GGCCAGACGACAACAAAGAGTCAACCCCACTGCCACGTCAAGAGCGAAGCGCCAC  
AGCTAAGACAAGAGGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGTT  
GACCATCTGTCTGCTTCTGTTTCCCCTTACTGCTCTTCCGATGGATGGAGATGAACCT  
GCAAACCGACCTGTCGAGCGTATGCAGGACAACATTTTCATCTGAGCAGTATCCCTTG  
TTTGAGAAGAGACGAGATTGTTGCACTCCGCCGAGGAAATGCAAAGACCGACGATG  
40 CAAACCCATGAAATGTTGCGCTGGACGATAACGTGTTGATGACCAACTTTATCACG  
GCTAGCTCAGTGTTTAGTGAATAAGTAAAATGATTGCAGTCTTGCTCAGATTGCTTT  
TGTGTTTTGGTCTAAGATCAATGACCAAACCGTTGTTTTGATGCGGATTGTCATATA  
TTTCTCGATTCCAATCCAACACTAGATGATTTAATCACGATAGATTAATTTTCTATCA  
ATGCCTTGATTTTTTCGTCTGTCATATCAGTTTTGTTTATATTTATTTTTCGTCACTGT  
45 CTACACAAACGCATGCATGCACGCATGCACGCACACACGCACGCACGCTCGCACAA  
ACATGCGCGCGCACGCACACACACACACACACAAACACACACACGAAGCAATC  
ACACAATTAGTTGACATTATTTATTTATTCATTGATGTATTTGTTATTTCGTTTGCTTGT

TTT TAGAATAGTTTGAGGCCGTCTTTTGGATTATTTGAACTGCTTTATTGTATACG  
AGTACTTCGTGCTTTGAAACACTGCTGAAAATAAAACAAACACTGACGTAGCAAAA  
AAAAAAA (SEQ ID NO:100)

5 **Translation:**

MMSKLGVLITICLLLFPLTALPMDGDEPANRPVERMQDNISSEQYPLFEKRRDCCTPPR  
KCKDRRCKPMKCCAGR (SEQ ID NO:101)

**Toxin Sequence:**

10 Arg-Asp-Cys-Cys-Thr-Xaa3-Xaa3-Arg-Lys-Cys-Lys-Asp-Arg-Arg-Cys-Lys-Xaa3-Met-Lys-  
Cys-Cys-Ala-# (SEQ ID NO:102)

**Name:**  $\mu$ -GIIC

15 **Species:** geographus

**Isolated:** Yes

**Toxin Sequence:**

20 Arg-Asp-Cys-Cys-Thr-Xaa3-Xaa3-Lys-Lys-Cys-Lys-Asp-Arg-Arg-Cys-Lys-Xaa3-Leu-Lys-  
Cys-Cys-Ala-# (SEQ ID NO:103)

**Name:** Gm3.1

25 **Species:** gloriamaris

**Cloned:** Yes

**DNA Sequence:**

CTCACTATAGGAATTCGAGCTCGGTACACGGGATCGATAGCAGTTCATGATGTCTAA  
30 ACTGGGAGCCTTGTTGACCATCTGTCTACTTCTGTTTTCCCTAACTGCTGTTCCGCTG  
GATGGAGATCAACATGCAGACCAACCTGCAGAGCGTCTGCATGACCGCCTTCCAAC  
TGAAAATCATCCCTTATATGATCCCGTCAAACGGTGTTGCGATGATTCGGAATGCGA  
CTATTCTTGCTGGCCTTGCTGTATGTTTGGATAACCTTTGTTATCGCGGCCTCGATAA  
GTGTCTAATGAATAAGTAAACGATTGCAGTAGGC (SEQ ID NO:104)

35

**Translation:**

MMSKLGALLTICLLLFSLTAVPLDGDQHADQPAERLHDRLPTENHPLYDPVKRCCDDSE  
CDYSCWPCCMFG (SEQ ID NO:105)

40 **Toxin Sequence:**

Cys-Cys-Asp-Asp-Ser-Xaa1-Cys-Asp-Xaa5-Ser-Cys-Xaa4-Xaa3-Cys-Cys-Met-Phe-# (SEQ ID  
NO:106)

45 **Name:** Gm3.2

**Species:** gloriamaris

**Cloned:** Yes



**DNA Sequence:**

5 GTTCATGATGTCTAAACTGGGAGTCTTGTTGATCATCTGTCTACTTCTGTTTCCCCTT  
ACTGCTGTTCCGCTGGATGGAGATCAACCTGCAGACCGATATGCAGAGCGTATGCA  
GGACGACATTTTCATCTGAACATCATCCCATGTTTGATGCCGTGAGAGGGTGTGCCA  
TCTGTTGGCATGCCGCTTCGGATGCTCGCCTTGTTGTTGGTGATCAGCTTTGTTATCG  
CGGCCTCATCAAGTGACTCTAATGCAA (SEQ ID NO:107)

**Translation:**

10 MMSKLGVLLIICLLLFPLTAVPLDGDQPADRYAERMQDDISSEHHPMFDAVRGCCHLLA  
CRFGCSPCCW (SEQ ID NO:108)

**Toxin Sequence:**

15 Gly-Cys-Cys-His-Leu-Leu-Ala-Cys-Arg-Phe-Gly-Cys-Ser-Xaa3-Cys-Cys-Xaa4<sup>^</sup> (SEQ ID  
NO:109)

**Name:** Gm3.3

**Species:** gloriamaris

20 **Cloned:** Yes

**DNA Sequence:**

25 GAGACGACAAGGAACAGTCAACCCACAGCCACGCCAAGAGCAGACAGCCACAGC  
TACGTGAAGAAGGGTGGAGAGAGGTTTCGTGATGTTGAAAATGGGAGTGGTGCTATT  
CATCTTCCTGGTACTGTTTCCCCTGGCAACGCTCCAGCTGGATGCAGATCAACCTGT  
AGAACGATATGCGGAGAAACAAACAGCTCCTCAACCCAGATGAAAGGAGGGAAATC  
ATATTGCATGCTCTGGGGACGCGATGCTGTTCTTGGGATGTGTGCGACCAACCCGAGT  
TGTACTTGCTGCGGCGGTTAGCGCCGAACATCCATGGCGCTGTGCTGGGCGGTTTAA  
TCCAACAACGACAGCGTTTGTGATTTTCATGTATCATTGCGCCACGTCTCTTGTCTA  
30 AGAATGACGAACATGATTGCACTCTGGTTCAGATTTCGTGTTCTTTTCTGACAATAA  
ATGACAAAACCTCAAAAAA (SEQ ID NO:110)

**Translation:**

35 MLKMGVVLFIPLVLFPLATLQLDADQPVERYAENKQLLNPDERREIILHALGTRCCSWD  
VCDHPSCTCCGG (SEQ ID NO:111)

**Toxin Sequence:**

40 Cys-Cys-Ser-Xaa4-Asp-Val-Cys-Asp-His-Xaa3-Ser-Cys-Thr-Cys-Cys-Gly-# (SEQ ID NO:112)

**Name:** La3.1

**Species:** laterculatus

**Cloned:** Yes

45 **DNA Sequence:**

CGACCTCAAGAAGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGTTGA  
CCATCTGTCTGCTTCTGTTTCCCCTTACTGCTCTCCGATGGATGGAGATCAACCTGC

AGACCGACCTGCAGAGCGTATGCAGGACGTTTCATCTGAACAGCATCCCTTGTATG  
ATCCCGTCAAACGGTGTTGCGACTGGCCATGCAGCGGATGCATCCCTTGTGCTAAT  
AGTAACAACGTGTTGATAACCAACTTTCTTACCACGACTACGTCAAGTGTCTAATGA  
ATAAGTAAAATGATTGCAGT (SEQ ID NO:113)

5

**Translation:**

MMSKLGVLITICLLLFPLTALPMDGDQPADRP AERMQDVSSEQHPLYDPVKRCCDWPC  
SGCIPCC (SEQ ID NO:114)

10 **Toxin Sequence:**

Cys-Cys-Asp-Xaa4-Xaa3-Cys-Ser-Gly-Cys-Ile-Xaa3-Cys-Cys-^ (SEQ ID NO:115)

15 **Name:** La3.2  
**Species:** laterculatus  
**Cloned:** Yes

**DNA Sequence:**

CGACCTCAAGAAGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGTGTA  
20 CCATCTGTCTGCTTCTGTTTCCCCCTTACTGCTCTGGATGGAGATCAACCTGCAGACC  
GACTTGCAGAGCGTATGCAGGACGACATTTTCATCTGAGCAGCATCCCTTTGAAAAG  
AGACGAGACTGTTGCACACCTCCGAAGAAATGCAGAGACCGACAATGCAAACCTGC  
ACGTTGTTGCGGAGGATAACGTGTTGATGACCAACTTTGTTATCACGGCTACGTCAA  
GTGTCTAGTGAATAAGTAAAACGATTGCAGT (SEQ ID NO:116)

25

**Translation:**

MMSKLGVLITICLLLFPLTALDGDQPADRLAERMQDDISSEQHPFEKRRDCCTPPKKCR  
DRQCKPARCCGG (SEQ ID NO:117)

30 **Toxin Sequence:**

Arg-Asp-Cys-Cys-Thr-Xaa3-Xaa3-Lys-Lys-Cys-Arg-Asp-Arg-Gln-Cys-Lys-Xaa3-Ala-Arg-  
Cys-Cys-Gly-# (SEQ ID NO:118)

35 **Name:** La3.3  
**Species:** laterculatus  
**Cloned:** Yes

**DNA Sequence:**

GGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGTGACCATCTGTCTGC  
40 TTCTGTTTCCCCCTTACTGCTCTTCCGATGGATGGAGATCAACTTGCACGCCGATCTGC  
AGAGCGTATGCAGGACAACATTTTCATCTGAGCAGCATCACCTCTTTGAAAAGAGAC  
GACCACCATGTTGCACCTATGACGGGAGTTGCCTAAAAGAATCATGCATGCGTAAA  
GCTTGTTGCGGATGATAACGTGTTGATGACCAACTTTGTTATCACGGCTACTCAAGT  
45 GTCTAATGAATAAGTAAAATGATTGCAGTA (SEQ ID NO:119)

**Translation:**

MMSKLGVLLTICLLLFPLTALPMDGDQLARRSAERMQDNISSEQHHLFEKRRPPCCTYD  
GSCLKESCMRKACCG (SEQ ID NO:120)

**Toxin Sequence:**

5 Arg-Xaa3-Xaa3-Cys-Cys-Thr-Xaa5-Asp-Gly-Ser-Cys-Leu-Lys-Xaa1-Ser-Cys-Met-Arg-Lys-  
Ala-Cys-Cys-# (SEQ ID NO:121)

10 **Name:** La3.3A  
**Species:** laterculatus  
**Cloned:** Yes

**DNA Sequence:**

GGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGTTGACCACCTGTCTGC  
15 TTCTGTTTCCCCTTACTGCTCTTCCGATGGATGGAGATCAACTTGCACGCCGACCTG  
CAGAGCGTATGCAGGACAACATTTTCATCTGAGCAGCATCCCTTCTTTGAAAGGAGA  
CGACCACCATGTTGCACCTATGACGGGAGTTGCCTAAAAGAATCATGCAAGCGTAA  
AGCTTGTTGCGGATAATAACGTGTTGATGACCAACTTTGTTATCACGGCTACTCAAG  
20 TGTCTAATGAATAAGTAAAATGATTGCAGTA (SEQ ID NO:122)

**Translation:**

MMSKLGVLLTICLLLFPLTALPMDGDQLARRPAERMQDNISSEQHPFFERRRPPCCTYD  
GSCLKESCKRKACCG (SEQ ID NO:123)

**Toxin Sequence:**

25 Arg-Xaa3-Xaa3-Cys-Cys-Thr-Xaa5-Asp-Gly-Ser-Cys-Leu-Lys-Xaa1-Ser-Cys-Lys-Arg-Lys-  
Ala-Cys-Cys-# (SEQ ID NO:124)

30 **Name:** Lp3.1  
**Species:** leopardus  
**Cloned:** Yes

**DNA Sequence:**

35 GGATCCATGATGTCTAAACTGGGAGTCTTGTTGACCGTCTGTCTGCTTCTGTTTCCCC  
TTACTGCTCTTCGGCTGGTTGGAGATCAACCTGCAGAGCGACCTGCAAAGCGTACGC  
AGGACGACATTCCAGATGGACAGCATCCGTAAATGATAGGCAGATAAACTGTTGC  
CCGTGGCCATGCCCTAGTACATGCCGCCATCAATGCTGCCATTAATGATAACGTGTT  
40 GATGACCAACTTTCTCGAG (SEQ ID NO:125)

**Translation:**

GSMMSKLGVLLTVCLLLFPLTALRLVGDQPAERPAKRTQDDIPDGQHPLNDRQINCCP  
WPCPSTCRHQCCH (SEQ ID NO:126)

**Toxin Sequence:**

45 Xaa2-Ile-Asn-Cys-Cys-Xaa3-Xaa4-Xaa3-Cys-Xaa3-Ser-Thr-Cys-Arg-His-Gln-Cys-Cys-His-^  
(SEQ ID NO:127)

5     **Name:**       Lv3.1  
      **Species:**   lividus  
      **Cloned:**    Yes

**DNA Sequence:**

GGATCCATGATGTCTAAACTGGGAGTCTTGTTGACCGTCTGTCTGCTTCTGTTTCCCC  
TTACTGCTCTTCGGCTGGTTAGAGATCAACCTGCAGAGCGACCTGCAAAGCGTACGC  
10   AGGACGACATTCCAAATGGACAGGATCCGTTAATTGATAGGCAGATAAATTGTTGC  
     CCTTGCCCATGCCCTGATTCATGCCACTATCAATGCTGCCACTGATAACGTGTTGAT  
     GACCAACTTTCTCGAG (SEQ ID NO:128)

**Translation:**

15   GSMMSKLGVLITVCLLLFPLTALRLVRDQPAERPAKRTQDDIPNGQDPLDRQINCCPW  
     PCPDSCHYQCCH (SEQ ID NO:129)

**Toxin Sequence:**

20   Xaa2-Ile-Asn-Cys-Cys-Xaa3-Xaa4-Xaa3-Cys-Xaa3-Asp-Ser-Cys-His-Xaa5-Gln-Cys-Cys-His-^  
     (SEQ ID NO:130)

25     **Name:**       L3.1  
      **Species:**   lynceus  
      **Cloned:**    Yes

**DNA Sequence:**

AAGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGTTGACCATCTGTCTG  
CTTCTGTTTCCCCTTACTGCTCTTCCGATGGATGGAGATCAATCTGCAGACCGACTTG  
30   CAGAGCGTATGCAGGACAACATTTTCATCTGAGCAGCATCCCTTCTTTGAAAAGAGA  
     GGACGAGACTGTTGCACACCTCCGAGGAAATGCAGAGACCGAGCCTGCAAACCTCA  
     ACGTTGTTGCGGAGGATAAGCTGTTGATGACCAACTTTGTTATACGGC (SEQ ID  
     NO:131)

**Translation:**

35   MMSKLGVLITICLLLFPLTALPMDGQDQSADRLAERMQDNISSEQHPFFEKRGRDCCTPP  
     RKCRDRACKPQRCCGG (SEQ ID NO:132)

**Toxin Sequence:**

40   Gly-Arg-Asp-Cys-Cys-Thr-Xaa3-Xaa3-Arg-Lys-Cys-Arg-Asp-Arg-Ala-Cys-Lys-Xaa3-Gln-  
     Arg-Cys-Cys-Gly-# (SEQ ID NO:133)

45     **Name:**       M3.1  
      **Species:**   magus  
      **Cloned:**    Yes

**DNA Sequence:**

CAAGAGGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGTTGACCATCTG  
TCTGCTTCTGTTTTCCCTTACTGCTCTTCCGATGGATGGAGATGAACCTGCAAACCG  
ACCTGTCGAGCGTATGCAGGACAACATTTTCATCTGAGCAGTATCCCTTGTTTGAGAA  
5 GAGACGAGATTGTTGCACTCCGCCGAAGAAATGCAAAGACCGACAATGCAAACCCC  
AGAGATGTTGCGCTGGACGATAACGTGTTGATGACCAACTTTATCACGGCTACGTCA  
AGTGTTTAGTGAATAAGTAAAATGATTGCAGTCTTGCTCAGATTTGCTTTTGTGTTTT  
GGTCTAAAGATCAATGACCAAACCGTTGTTTTGATGCGGATTGTCATATAATTTCTCG  
ATTCCAATCCAACACTAGATGATTTAATCACGATAGATTAATTTTCTATCAATGCCT  
10 TGATTTTTCTGCTCTGTCATATCAGTTTTGTTTATATTTATTTTTTCGTCAGTGTCTACAC  
AAACGCATGCATGCACGCATGCACGCACACACGCACGCACGCTCGCACAAACATGC  
GCGCGCACGCACACACACACACACACACAAACACACACACGAAGCAATCACAC  
AATTAGTTGACATTATTTATTTATTCATTGATGATTTGTTATTCGTTTGCTTTGTTTTT  
AGAATAGTTTGAGGCCGTCTTTTTGGATTATTTGAACTGCTTTATTGTATACGAGTA  
15 CTTCGTGCGGGGAAACACTGCTGAAAATAAAACAAACACTGACGTAGCAAAAAAA  
AAAAA (SEQ ID NO:134)

**Translation:**

MMSKLGVLITICLLLFPLTALPMDGDEPANRPVERMQDNISSEQYPLFEKRRDCCTPPK  
20 KCKDRQCKPQRCCAGR (SEQ ID NO:135)

**Toxin Sequence:**

Arg-Asp-Cys-Cys-Thr-Xaa3-Xaa3-Lys-Lys-Cys-Lys-Asp-Arg-Gln-Cys-Lys-Xaa3-Gln-Arg-  
Cys-Cys-Ala-# (SEQ ID NO:136)  
25

Name: M3.2  
Species: magus  
Cloned: Yes  
30

**DNA Sequence:**

CAAGAGGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGTTGACCATCTG  
TCTGCTTCTGTTTTCCCTTACTGCTCTTCCAATGGATGGAGATCAACCTGCAGACCA  
ACCTGCAGATCGTATGCAGGACGACATTTTCATCTGAGCAGTATCCCTTGTTTGATAT  
35 GAGAAAAAGGTGTTGCGGCCCCGGCGGTTTCATGCCCCGTATATTTTCAGAGACAATT  
TTATTTGTGGTTGTTGTTAAATGACAACGTGTCGATGACCAACTTCATTATCACGAC  
TACGCCAAGTGTCTAATGAATAAAATAAAATGATTGCAGTCTCGCTCAGATTTGCTTT  
TGTATTTTGGTCTAAAGATCAATGACCAAACCGTTGTTTTGGTGTGGATTTTCATATA  
TTTCTCGAGTCCTATCCAACACTAGATGATTTAATCACGATAGATCTGATTTTTTTAT  
40 CAAAGGCTTGGTTTTTCGCTCTGTACATCAGTTTTGTTTATATTTAATTTTCGTCAGT  
GATTACACACACGCATGAACGCACAGAGTACTAACACATACACACACACACACACAC  
CACACACACACACACACACACACACACACACACACACACGCGCGCGCGCGGCG  
CCATCTAGTAGCGCCGCGACGACACACAC (SEQ ID NO:137)

**Translation:**

MMSKLGVLITICLLLFPLTALPMDGDQPADQPADRMQDDISSEQYPLFDMRKRC CGPG  
GSCP VYFRDNFICGCC (SEQ ID NO:138)  
45

**Toxin Sequence:**

Cys-Cys-Gly-Xaa3-Gly-Gly-Ser-Cys-Xaa3-Val-Xaa5-Phe-Arg-Asn-Phe-Ile-Cys-Gly-Cys-  
Cys-^ (SEQ ID NO:139)

5

**Name:** M3.3  
**Species:** magus  
**Cloned:** Yes

10

**DNA Sequence:**

CAAGAGGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGTTGACCATCTG  
TTTGCTTCTGTTTCCCTTACTGCTCTTCCGAGGGATGGAGATCAATCTGTAGACCGA  
CCTGCAGAGCGTATGCAGGACGACATTTTCATCTGAGCTGCATCCCTTGTCAATCAGA  
15 AAAAGAATGTGTTGCGGCGAGAGTGCGCCATGCCCCAGCTATTTCAGAAACAGTCA  
GATTTGTCATTGTTGTTAAATGACAACGTGTCGATGACCACCTTCGTTATCACGACT  
AATGATAAGTAAAATGATTGCAGTCTCGCTCAGATTTGCTTTTGTATTTTGGTCTAA  
AGATCAATGACCAAACCGTTGTTTTGATGTGGATTTTCATATATTTCTCGAGTCCTAT  
CCAACACTAGATGATTTAATCACGATAGATCTGATTTTTTTATCAAAGCCTTGTTTT  
20 TCGTCTGTACATCAGTTTTGTTTATATTTAATTTTCGTCACTGATTACACACACGC  
ATGAACGCACAGACGTACTAACACATACACACACACACACACACACACACACAC  
ACACACACACACACACACAC (SEQ ID NO:140)

**Translation:**

25 MMSKLGVLITICLLFPLTALPRDGDQSVDRPAERMQDDISSELHPLSIRKRMCCGESAP  
CPYFRNSQICHCC (SEQ ID NO:141)

**Toxin Sequence:**

30 Met-Cys-Cys-Gly-Xaa1-Ser-Ala-Xaa3-Cys-Xaa3-Ser-Xaa5-Phe-Arg-Asn-Ser-Gln-Ile-Cys-His-  
Cys-Cys-^ (SEQ ID NO:142)

**Name:** M3.4  
**Species:** magus  
35 **Cloned:** Yes

**DNA Sequence:**

CAAGAGGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGTTGACCATCTG  
TCTGCTTCTGTTTCCCTTACTGCTCTTCCAATGGATGGAGATCAACCTGCAGACCA  
40 ACCTGCAGATCGTATGCAGGACGACATTTTCATCTGAGCAGTATCCCTTGTGTTGATAA  
GAGACAAAAGTGTTGCGGCCCCGGCGGTTTCATGCCCCGTATATTTACAGACAATTT  
TATTTGTGGTTGTTGTTAAATGACAACGTGTCGATGACCAACTTCATTATCACGACT  
ACGCCAAGTGTCTAATGAATAAATAAAATGATTGCAGTCTCGCTCAGATTTGCTTTT  
GTATTTGGTCTAAAGATCAATGACCAAACCGTTGTTTTGGTGCTGGATTTTCATATA  
45 TTTCTCGATTCCATCCAACACTAGATGATTTAATCACGATAGATCTGATTTTTTTAT  
CAATGCCTTAATTTTTTGCTCTGTCATATCAGTTTTGTTTATAT (SEQ ID NO:143)

**Translation:**

MMSKLGVLLTICLLLFPLTALPMDGDQPADQPADRMQDDISSEQYPLFDKRQKCCGPG  
GSCP VYFTDNFICGCC (SEQ ID NO:144)

5 **Toxin Sequence:**

Xaa2-Lys-Cys-Cys-Gly-Xaa3-Gly-Gly-Ser-Cys-Xaa3-Val-Xaa5-Phe-Thr-Asp-Asn-Phe-Ile-Cys-  
Gly-Cys-Cys-^ (SEQ ID NO:145)

10 **Name:** M3.5  
**Species:** magus  
**Cloned:** Yes

**DNA Sequence:**

15 CAAGAGGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGTTGACCATCTG  
TCTGCTTCTGTTTCCCTTACTGCTCTTCCAATGGATGGAGATCAACCTGCAGACCA  
ACCTGCAGATCGTATGCAGGACGACATTTTCATCTGAGCAGTATCCCTTGTTTGATAA  
GAGACAAAAGTGTTGCGGCCCCGGCGGTTTCATGCCCCGTATATTCAGAGACAATTT  
TATTTGTGGTTGTTGTTAAATGACAACGTGTCGATGACCATCTTCATTATCACGACT  
20 ACGCCAAGTGTCTAATGAATAAAATAAAATGATTGCAGTCTCGCTCAGATTTGCTTTT  
GTATTTTGGTCTAAAGATCAATGACCAAACCGTTGTTTTGGTGTGGATTTTCATATAT  
TTCTCGATTCCCTATCCAACACTAGATGATTTAATCACGATAGATCTGATTTTTT (SEQ  
ID NO:146)

25 **Translation:**

MMSKLGVLLTICLLLFPLTALPMDGDQPADQPADRMQDDISSEQYPLFDKRQKCCGPG  
GSCP VYFRDNFICGCC (SEQ ID NO:147)

**Toxin Sequence:**

30 Xaa2-Lys-Cys-Cys-Gly-Xaa3-Gly-Gly-Ser-Cys-Xaa3-Val-Xaa5-Phe-Arg-Asp-Asn-Phe-Ile-  
Cys-Gly-Cys-Cys-^ (SEQ ID NO:148)

35 **Name:** U001  
**Species:** magus  
**Isolated:** No

**Toxin Sequence:**

40 Xaa2-Lys-Cys-Cys-Ser-Gly-Gly-Ser-Cys-Xaa3-Leu-Xaa5-Phe-Arg-Asp-Arg-Leu-Ile-Cys-Xaa3-  
Cys-Cys-^ (SEQ ID NO:149)

45 **Name:** Comatose/Death  
**Species:** marmoreus  
**Isolated:** Yes

**Toxin Sequence:**

Ser-Lys-Gln-Cys-Cys-His-Leu-Ala-Ala-Cys-Arg-Phe-Gly-Cys-Thr-Xaa3-Cys-Cys-Asn-^ (SEQ ID NO:150)

5   **Name:**       Mr3.1  
     **Species:**   marmoreus  
     **Cloned:**    Yes

**DNA Sequence:**

10   CAAGAAGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGTTGACCATCTG  
     TCTGCTTCTGTTTCCCGTTACTGCTCTTCCGATGGATGGTGATCAACCTGCAGACCGA  
     CTTGTAAGCGTATGCAGGACAACATTTTCATCTGAGCAGCATCCCTTCTTTGAAAAG  
     AGAAGAGGAGGCTGTTGCACACCTCCGAGGAAATGCAAAGACCGAGCCTGCAAAC  
     CTGCACGTTGCTGCGGCCAGGATAACGTGTTGATGACCAACTTTGTTATCACGGCT  
15   ACGTCAAGTGTCTAGTGAATAAGTAAAACGATTGCAG (SEQ ID NO:151)

**Translation:**

     MMSKLGVLITICLLLPVTALPMDGDQPADRLVERMQDNISSEQHPFFEKRRGGCCTPP  
     RKCKDRACKPARCCGPG (SEQ ID NO:152)

20

**Toxin Sequence:**

     Arg-Gly-Gly-Cys-Cys-Thr-Xaa3-Xaa3-Arg-Lys-Cys-Lys-Asp-Arg-Ala-Cys-Lys-Xaa3-Ala-Arg-  
     Cys-Cys-Gly-Xaa3-# (SEQ ID NO:153)

25

**Name:**       Mr3.2  
     **Species:**   marmoreus  
     **Cloned:**    Yes

30   **DNA Sequence:**

     GAGCTCGGTACCCCGACCTCAAGAGGGATCGATAGCAGTTCATGATGTCTAAACTG  
     GGAATCTTGTTGACCATCTGTCTACTTCTATTTCCCCTTACTGCTGTTCCGCTGGATG  
     GAGATCAACCTGCAGACCGACCTGCAGAGCGTATGCAGGACGACATTTTCATCTGAA  
     CATCATCCCTTTTTTGTATCCCGTCAAACGGTGTTGCAGGTTATCATGCGGCCTGGGA  
35   TGCCACCCTTGTTGTGGATGACCAGCTTTGTTATCGCGGCCTCATCAAGTGTCTAAT  
     GAATAAGTAAAA (SEQ ID NO:154)

**Translation:**

     MMSKLGILLITICLLLFPLTAVPLDGDQPADRPAERMQDDISSEHHPFFDPVKRCCRLSCG  
40   LGCHPCCG (SEQ ID NO:155)

**Toxin Sequence:**

     Cys-Cys-Arg-Leu-Ser-Cys-Gly-Leu-Gly-Cys-His-Xaa3-Cys-Cys-# (SEQ ID NO:156)

45

**Name:**       Mr3.3  
     **Species:**   marmoreus



Cloned: Yes

**DNA Sequence:**

GGCCTACACCAAGCTTGCATGCCTGCAGGTCGACTCTAGAGGATCCCCGATCGATA  
5 GCAGTTCATGATGTCTAGACTGGGAGTCTTGTTGACCATCTGTCTACTTCTGTTTCCC  
CTTACTGCTGTTCCGCTGGATGGAGATCAACCTGCGGACCGACCTGCAGAGCGCCTG  
CAGGACGACATTTTCATCTGAACATCATCCCCATTTTGATTCCGGCAGAGAGTGTTGC  
GGTTCGTTTCGCATGCCGCTTTGGATGCGTGCTTGTTGTGTATGACCAGCTTTGTTAT  
CACGGCCTCATCGAGTGTCTAATGAATAAGTAAAACGATTGCAGTAGGCGGGTACC  
10 GAGCTCGAATTCC (SEQ ID NO:157)

**Translation:**

MMSRLGVLLTICLLLFPLTAVPLDGDQPADRPAERLQDDISSEHHPHFDSGRECCGSFAC  
15 RFGCVPCCV (SEQ ID NO:158)

**Toxin Sequence:**

Xaa1-Cys-Cys-Gly-Ser-Phe-Ala-Cys-Arg-Phe-Gly-Cys-Val-Xaa3-Cys-Cys-Val-^ (SEQ ID  
20 NO:159)

Name: Mr3.4  
Species: marmoreus  
Cloned: Yes

**DNA Sequence:**

CGACCTCAAGAGGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGTTGA  
25 CCATCTGTCTACTTCTATTTCCCCTTACTGCTGTTCCGCTGGATGGAGACCAACCTGC  
AGACCGACCTGCAGAGCGTATGCAGGACGACATTTTCATCTGAACGTCATCCTTTTTT  
TGATCGCAGCAAACAGTGTTGCCATCTGCCGGCATGCCGCTTCGGATGTACGCCTTG  
30 TTGTTGGTGATCAGCTTTGTTATCGCGTCCTCATCAAGTGTCTAATGAATAAGTAAA  
ATGATTGCAG (SEQ ID NO:160)

**Translation:**

MMSKLGVLLTICLLLFPLTAVPLDGDQPADRPAERMQDDISSERHPFFDRSKQCCHLPA  
35 CRFGCTPCCW (SEQ ID NO:161)

**Toxin Sequence:**

Ser-Lys-Gln-Cys-Cys-His-Leu-Xaa3-Ala-Cys-Arg-Phe-Gly-Cys-Thr-Xaa3-Cys-Cys-Xaa4-^  
40 (SEQ ID NO:162)

Name: Mr3.5  
Species: marmoreus  
Cloned: Yes

**DNA Sequence:**

GGATCCATGATGTCTAAACTGGGAGTCTTGTTGACCATCTGTCTGCTTCTGTTTCCCC

TTACTGCTCTTCCGCTGGATGGAGATCAACCTGCAGACCAACGTGCAGAGCGTACG  
CAGGCCGAGAAGCATTCTTGCCTGATCCGAGAATGGGCTGTTGCCCGTTTCCATGC  
AAAACCAGTTGCACTACTTTGTGTTGCGGGTGATGATAACGTGTTGATGACCAACTT  
TCTCGAG (SEQ ID NO:163)

5

**Translation:**

GSMMSKLGVLITICLLFPLTALPLDGDQPADQRAERTQAEKHSLPDRMGCCPFCKT  
SCTTLCCG (SEQ ID NO:164)

10 **Toxin Sequence:**

Met-Gly-Cys-Cys-Xaa3-Phe-Xaa3-Cys-Lys-Thr-Ser-Cys-Thr-Thr-Leu-Cys-Cys-# (SEQ ID  
NO:165)

15 **Name:** U014  
**Species:** marmoreus  
**Isolated:** Yes

**Toxin Sequence:**

20 Cys-Cys-His-Xaa4-Asn-Xaa4-Cys-Asp-His-Leu-Cys-Ser-Cys-Cys-Gly-Ser-^ (SEQ ID NO:166)

**Name:** U017  
**Species:** marmoreus  
25 **Cloned:** Yes

**DNA Sequence:**

GCCAAGCTTGCATGCCTGCAGGATGACTCTAGAGGATCCCCACCTCAAGAGGGATC  
GATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGTTGACCATCTGTCTACTTCTGTT  
30 TGCCCTTACTGCTGTTCCGCTGGATGGAGATCAACCTGCAGACCGACCTGCAGAACG  
TATGCAGGACGACATTTTCATCTGAACGTCATCCCATGTTTGATGCCGTCAGAGATTG  
TTGCCCGTTGCCGGCATGCCCTTTGGATGCAACCCTTGTTGTGGATGACCAGCTTT  
GTTATCGGGACCTCATCAAGTGTCTAATGAATAAGTAAAAAACGATTCGAGTGGGT  
ACCGAGCTCGAATTCC (SEQ ID NO:167)

35

**Translation:**

MMSKLGVLITICLLFALTAVPLDGDQPADRPAERMQDDISSERHPMFDAVRDCCPLP  
ACPFGCNPCCG (SEQ ID NO:168)

40 **Toxin Sequence:**

Asp-Cys-Cys-Xaa3-Leu-Xaa3-Ala-Cys-Xaa3-Phe-Gly-Cys-Asn-Xaa3-Cys-Cys-# (SEQ ID  
NO:169)

45 **Name:** U019  
**Species:** marmoreus  
**Isolated:** Yes

**Toxin Sequence:**

Cys-Cys-Ala-Xaa3-Ser-Ala-Cys-Arg-Leu-Gly-Cys-Arg-Xaa3-Cys-Cys-Arg-^ (SEQ ID NO:170)

5

**Name:** U020  
**Species:** marmoreus  
**Isolated:** Yes

10 **Toxin Sequence:**

Cys-Cys-Ala-Xaa3-Ser-Ala-Cys-Arg-Leu-Gly-Cys-Arg-Xaa3-Cys-Cys-Arg-^ (SEQ ID NO:171)

15 **Name:** U022  
**Species:** marmoreus  
**Isolated:** Yes

**Toxin Sequence:**

Cys-Cys-Ala-Xaa3-Ser-Ala-Cys-Arg-Leu-Gly-Cys-Arg-Xaa3-Cys-Cys-Arg-^ (SEQ ID NO:172)

20

**Name:** U024  
**Species:** marmoreus  
**Isolated:** Yes

25

**Toxin Sequence:**

Gly-Cys-Cys-Gly-Ser-Phe-Ala-Cys-Arg-Phe-Gly-Cys-Val-Xaa3-Cys-Cys-Val-^ (SEQ ID NO:173)

30

**Name:** Nb3.1  
**Species:** nobilis  
**Cloned:** Yes

35 **DNA Sequence:**

GGATCCATGATGTCTAAACTGGGAGTCTTGTTGACCATCTGTCTACTTCTGTTTCCCC  
TTACTGCTCTTCCGCTGGATGAAGATCAACCGGTACACCGACCTGCAGAGCGTATGC  
AGGACATTTTCATCTGATCAACATCTCTTCTTTGATCTCATCAAACGGTGCTGCGAGT  
TGCCATGCGGGCCAGGCTTTTGCGTCCCTTGTTGCTGACATCAATAACGTGTTGATG  
40 ACCAACTTTCTCGAG (SEQ ID NO:174)

**Translation:**

GSMMSKLGVLITICLLFPLTALPLDEDQPVHRPAERMQDISSDQHLFFDLIKRCCELPC  
GPGFCVPCC (SEQ ID NO:175)

45

**Toxin Sequence:**

Cys-Cys-Xaa1-Leu-Xaa3-Cys-Gly-Xaa3-Gly-Phe-Cys-Val-Xaa3-Cys-Cys-^ (SEQ ID NO:176)

Name: Nb3.2  
Species: nobilis  
5 Cloned: Yes

**DNA Sequence:**

GGATCCATGATGTCTAAACTGGGAGTCTTGTTGACCATCTGTCTACTTCTGTTTCCCC  
TTACTGCTTTTCCGATGGATGGAGATCAACCTGCAGACCAACCTGCAGATCGTATGC  
10 AGGACGACATTTTCATCTGAGCAGTATCCCTTGTTTGATAAGAGACAAAAGTGTTGCA  
CTGGGAAGAAGGGGTCATGCTCCGGCAAAGCATGCAAAAATCTCAAATGTTGCTCT  
GGACGATAACGTGTTGATGACCAACTTTCTCGAG (SEQ ID NO:177)

**Translation:**

15 GSMMSKLGVLLTICLLFPLTAFPMGDGQPADQPADRMQDDISSEQYPLFDKRQKCC<sup>T</sup>  
GKKGSCSGKACKNLKCCSGR (SEQ ID NO:178)

**Toxin Sequence:**

Xaa2-Lys-Cys-Cys-Thr-Gly-Lys-Lys-Gly-Ser-Cys-Ser-Gly-Lys-Ala-Cys-Lys-Asn-Leu-Lys-  
20 Cys-Cys-Ser-# (SEQ ID NO:179)

Name: Pu3.1  
Species: pulicarius  
25 Cloned: Yes

**DNA Sequence:**

GGATCCATGATGTCTAAACTGGGAGTTTTGTTGACCATCTGTCTGCTTCTGTTTCCCC  
TTACTGCTGTTCCGCTGGATGGAGATCAACCTGCAGACCGACCTGCAGAGCGTATGC  
30 AGGACATTGCAACTGAACAGCATCCCTTCTTTGATCCCGTCAAACGGTGTTGCAACA  
GCTGTTACATGGGATGCATCCCTTGTTGCTTCTAGTAATAACGTGTTGATGACCAAC  
TTTCTCGAG (SEQ ID NO:180)

**Translation:**

35 GSMMSKLGVLLTICLLFPLTAVPLDGDQPADRPAERMQDIATEQHPFDPVKRCCNSC  
YMGCI<sup>PCCF</sup> (SEQ ID NO:181)

**Toxin Sequence:**

Cys-Cys-Asn-Ser-Cys-Xaa5-Met-Gly-Cys-Ile-Xaa3-Cys-Cys-Phe-<sup>^</sup> (SEQ ID NO:182)  
40

Name: Qc3.1  
Species: quercinus  
45 Cloned: Yes

**DNA Sequence:**

GGATCCATGATGTCTAAACTGGGAGTCTTGTTGACCATCTGTCTGCTTCTGTTTCCCC

TTACAGCTCTTCAGCTGGATGGAGATCAACCTGCAGACCGACCTGCAGAGCGTACG  
CAGGACATTGCATCTGAACAGTATCGAAAGTTTGATCAGAGACAGAGGTGTTGCCA  
GTGGCCATGCCCCGGTAGTTGCAGATGCTGCCGTA CTGGTTAACGTGTTGATGACCA  
ACTTTCTCGAG (SEQ ID NO:183)

5

**Translation:**

GSMMSKLGVLITICLLLFPLTALQLDGDQPADRPAERTQDIASEQYRKFDQRQRCCQW  
PCPGSCRCRTG (SEQ ID NO:184)

10 **Toxin Sequence:**

Xaa2-Arg-Cys-Cys-Gln-Xaa4-Xaa3-Cys-Xaa3-Gly-Ser-Cys-Arg-Cys-Cys-Arg-Thr-# (SEQ ID  
NO:185)

15 **Name:** QcIIIA  
**Species:** quercinus  
**Isolated:** Yes

**Toxin Sequence:**

20 Cys-Cys-Ser-Gln-Asp-Cys-Leu-Val-Cys-Ile-Xaa3-Cys-Cys-Xaa3-Asn-# (SEQ ID NO:186)

**Name:** QcIIIB  
**Species:** quercinus  
25 **Isolated:** Yes

**Toxin Sequence:**

Cys-Cys-Ser-Arg-His-Cys-Xaa4-Val-Cys-Ile-Xaa3-Cys-Cys-Xaa3-Asn-? (SEQ ID NO:187)

30

**Name:** R3.1  
**Species:** radiatus  
**Isolated:** Yes  
**Cloned:** Yes

35

**DNA Sequence:**

TCAAGAAGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGTTGACCATCT  
GTCTGCTTCTGTTTCCCCTTACTGCTCTTCCGATGGATGGAGATCAACCTGTAGACCG  
ACTTGCAGAGCGTATGCAGGACAACATTTTCATCTGAGCAGCATACCTTCTTTGAAAA  
40 GAGACTACCATCGTGTGCTCCCTTAACTTGCGGCTTTGCCAGTACCAGCATGCAA  
ACGTAACCCTTGTTGCACAGGATAACGTGTTGATGACCAACTTTGTTATCACGGCTA  
CGTCAAGTGTCTAGTGAATAAGTAAAACGATTGCAGT (SEQ ID NO:188)

**Translation:**

45 MMSKLGVLITICLLLFPLTALPMDGDQPVDRLAERMQDNISSEQHTFFEKRLPSCCSLN  
LRLCPVPA CKRNPCCTG (SEQ ID NO:189)

**Toxin Sequence:**

Leu-Xaa3-Ser-Cys-Cys-Ser-Leu-Asn-Leu-Arg-Leu-Cys-Xaa3-Val-Xaa3-Ala-Cys-Lys-Arg-Asn-Xaa3-Cys-Cys-Thr-# (SEQ ID NO:190)

5

**Name:** R3.2  
**Species:** radiatus  
**Cloned:** Yes

10 **DNA Sequence:**

AGGTCGACTCTAGAGGATCCCCAAGGATCGATAGCAGTTCATGATGTCTAAACTGG  
GAGTCTTGTTGACCATCTGTCTGCTTCTGTTCCCTTACTGCTCTCCGATGGATGG  
AGATCAACCTGCAGACCGACTTGCAGAGCGTATGCAGGACGACATTTTCATCTGAGC  
AGCATCCCTTCTTTAAAAAGAGACAACAAAGATGTTGCACCGTTAAGAGGATTTGT  
15 CCAGTACCAGCATGCAGAAGTAAACCTTGTGCAAATCATAACGTATTGATGACCA  
ACTTTGTTATCACGGCTACGTCAAGTGTCTAGTGAATAAGTAAAATGATTGCAG  
(SEQ ID NO:191)

**Translation:**

20 MMSKLGVLITICLLFPLTALPMDGDQPADRLAERMQDDISSEQHPFFKKRQRCCTV  
KRICPVPACRSKPCKS (SEQ ID NO:192)

**Toxin Sequence:**

25 Xaa2-Gln-Arg-Cys-Cys-Thr-Val-Lys-Arg-Ile-Cys-Xaa3-Val-Xaa3-Ala-Cys-Arg-Ser-Lys-Xaa3-  
Cys-Cys-Lys-Ser-^ (SEQ ID NO:193)

**Name:** R3.3  
**Species:** radiatus  
30 **Cloned:** Yes

**DNA Sequence:**

ACCTCAAGAAGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGTTGACC  
ATCTGTCTGCTTCTGTTCCCGTTACTGCTCTCCGATGGATGGTGATCAACCTGCAG  
35 ACCGACTTGTAGAGCGTATGCAGGACAACATTTTCATCTGAGCAGCATCCCTTCTTTG  
AAAAGAGAAGAGGAGGCTGTTGCACACCTCCGAGGAAATGCAAAGACCGAGCCTG  
CAAACCTGCACGTTGCTGCGGCCAGGATAACGTGTTGATGACCAACTTTGTTATCA  
CGGCTACGTCAAGTGTCTAGTGAATAAGTAAAACGATTGCAGT (SEQ ID NO:194)

40 **Translation:**

MMSKLGVLITICLLFVPTALPMDGDQPADRLVERMQDNISSEQHPFFEKRRGGCCTPP  
RKCKDRACKPARCCGPG (SEQ ID NO:195)

**Toxin Sequence:**

45 Arg-Gly-Gly-Cys-Cys-Thr-Xaa3-Xaa3-Arg-Lys-Cys-Lys-Asp-Arg-Ala-Cys-Lys-Xaa3-Ala-Arg-  
Cys-Cys-Gly-Xaa3-# (SEQ ID NO:196)

**Name:** Ra3.1  
**Species:** rattus  
5 **Cloned:** Yes

**DNA Sequence:**

GGATCCATGATGTCTAAACTGGGAGTCTTGGTGACCATCTGCCTGCTTCTGTTCCCT  
CTTGCTGCTTTTCCACTGGATGGAGATCAACCTGCAGACCACCCTGCAAAGCGTACG  
10 CAAGATGACAGTTCAGCTGCCCTGATCAATGCCTGGCTTGATGAATCCCAGACTTGC  
TGCAGTAACTGCGGTGAAGATTGTGATGGTTGTTGCCAGTAACGTGTTGATGACCAA  
CTTTCTCGAG (SEQ ID NO:197)

**Translation:**

15 GSMMSKLGVLVTICLLFPLA AFPLDGDQPADHPAKRTQDDSSAALINAWLDESQTCCS  
NCGEDCDGCCQ (SEQ ID NO:198)

**Toxin Sequence:**

20 Xaa2-Thr-Cys-Cys-Ser-Asn-Cys-Gly-Xaa1-Asp-Cys-Asp-Gly-Cys-Cys-Gln-^ (SEQ ID  
NO:199)

**Name:** Sm3.1  
**Species:** stercusmuscarum  
25 **Cloned:** Yes

**DNA Sequence:**

GACCTCAAGAGGGATCGATAGCAGTTCGTGATGTCTAAACTGGGAGTCTTGTGAC  
CATCTGTCTGCTTCTGTTTCCTCTTACTGCTCTTCCGATGGATGGAGATCAACCTGCA  
30 GACCAACCTGCAGATCGTATGCAGGACGACATTTTCATCTGAGCAGTATCCCTTGTTT  
GATAAGAGACAAAAGTGTGCACTGGGAAGAAGGGGTCATGCTCCGGCAAAGCAT  
GCAAAAATCTCAAATGTTGCTCTGGACGATAACGTGTTGATGACCAACTTTGTTATC  
ACGGCTACGTCAAAGTGTCTAATGAATAAGTAAAACGATTGCAGT (SEQ ID NO:200)

**Translation:**

35 MSKLGVLLTICLLFPLTALPMDGDQPADQPADRMQDDISSEQYPLFDKRQKCCTGKK  
GSCSGKACKNLKCCSGR (SEQ ID NO:201)

**Toxin Sequence:**

40 Xaa2-Lys-Cys-Cys-Thr-Gly-Lys-Lys-Gly-Ser-Cys-Ser-Gly-Lys-Ala-Cys-Lys-Asn-Leu-Lys-  
Cys-Cys-Ser-# (SEQ ID NO:202)

**Name:** U034  
45 **Species:** stercusmuscarum  
**Isolated:** Yes  
**Cloned:** Yes

**DNA Sequence:**

5 GATCGATAGCAGTTCGTGATGTCTAAACTGGGAGTCTTGTTGACCATCTGTCTGCTT  
CTGTTTCCCCTTACTGCTCTTCCGATGGATGGAGATCAACCTGCAGACCAACCTGCA  
GATCGTATGCAGAACGACATTTTCATCTGAGCAGTATCCCTTGTTTGATAAGAGACAA  
AAGTGTGCGGCCCCGGCGCGTCATGCCCCAGATATTTCAAAGACAATTTTATTTGT  
GGTTGTTGTTAAATGACAACGTGTCGATGACCAACTTCGTTATCACGACTTCGCCAA  
GTGTCTAATGAATAAGTAAAACGATTGCAGT (SEQ ID NO:203)

**Translation:**

10 MSKLGVLLTICLLFPLTALPMDGDQPADQPADRMQNDISSEQYPLFDKRQKCCGPGAS  
CPRYFKDNFICGCC (SEQ ID NO:204)

**Toxin Sequence:**

15 Xaa2-Lys-Cys-Cys-Gly-Xaa3-Gly-Ala-Ser-Cys-Xaa3-Arg-Xaa5-Phe-Lys-Asp-Asn-Phe-Ile-  
Cys-Gly-Cys-Cys-^ (SEQ ID NO:205)

**Name:** S3.1

20 **Species:** striatus

**Cloned:** Yes

**DNA Sequence:**

25 CGACCTTTCAAGAGGGATCGATAGCAGTTCGCGATGTCTAAACTGGGGGTATTGTTG  
ACCATCTGTCTGCTTCTGTTTCCCCTTACTGCTCTTCCGATGGATGAAGATCAACCTG  
CAGACCAACTTGAAGATCGTATGCAGGACGACATTTTCATCTGAGCAGTATCCCTCGT  
TTGTTAGGAGACAAAAGTGTGCGGCGAAGGCTCGTCATGCCCCAAATATTTCAA  
AACAATTTTATTTGTGGTTGTTGTTAAATGACAACGTGTCGATGACCAACTTCGTTA  
TCACGACTACGCCAAGTGTCTTGTCTAATGATAATAAAATGATTCC (SEQ ID NO:206)

30

**Translation:**

MSKLGVLLTICLLFPLTALPMDEDQPADQLEDQMDDISSEQYPSFVRRQKCCGEGSS  
CPKYFKNNFICGCC (SEQ ID NO:207)

**Toxin Sequence:**

35 Xaa2-Lys-Cys-Cys-Gly-Xaa1-Gly-Ser-Ser-Cys-Xaa3-Lys-Xaa5-Phe-Lys-Asn-Asn-Phe-Ile-Cys-  
Gly-Cys-Cys-^ (SEQ ID NO:208)

40 **Name:** S3.2

**Species:** striatus

**Cloned:** Yes

**DNA Sequence:**

45 GGATCCATGATGTCTAAACTGGGAGTCTTGTTGACCGTCTGTCTGCTTCTGTTTCCCC  
TTACTGCTCTTCCGCTGGATGGAGATCAACCTGCAGACCGACCTGCAGAGCGTATGC  
AGGACGACATTTTCATCTGACGAGCATCCCTTGTTTGATAAGAGACAAAACCTGTTGCA



56

ATGGGGGATGCTCCAGCAAATGGTGCAGAGATCACGCACGTTGTTGCGGTCGATGA  
TAACGTGTTGATGACCAACTTTCTCGAG (SEQ ID NO:209)

**Translation:**

5 GSMMSKLGVLLTVCLLLFPLTALPLDGDQPADRP AERMQDDISSDEHPLFDKRQNCCN  
GGCSSKWCRDHARCCGR (SEQ ID NO:210)

**Toxin Sequence:**

10 Xaa2-Asn-Cys-Cys-Asn-Gly-Gly-Cys-Ser-Ser-Lys-Xaa4-Cys-Arg-Asp-His-Ala-Arg-Cys-Cys-#  
(SEQ ID NO:211)

Name: Ts3.1

Species: tessulatus

15 Cloned: Yes

**DNA Sequence:**

GGATCCATGATGTCTAAACTGGGAGTCTTGTTGACCATGTGTCTGCTTCTGTTTCCCC  
TTACTGCTGTTCCGCTGGATGGAGATCAACCTGCAGACCGACCTGCAGAGCGTAGG  
20 CAGGACATTGCAACTGACGATCATCCTTTGTTGATCCCGTCAAACGGTGCTGCCAC  
AAATGCTATATGGGATGCATCCCTTGTTGCATTTAGTAACGTGTTGATGACCAACTT  
TCTCGAG (SEQ ID NO:212)

**Translation:**

25 GSMMSKLGVLLTMCLLLFPLTAVPLDGDQPADRP AERRQDIATDDHPLFDPVKRCCHK  
CYMGCI PCCI (SEQ ID NO:213)

**Toxin Sequence:**

30 Cys-Cys-His-Lys-Cys-Xaa5-Met-Gly-Cys-Ile-Xaa3-Cys-Cys-Ile-^ (SEQ ID NO:214)

Name: Ts3.2

Species: tessulatus

35 Cloned: Yes

**DNA Sequence:**

GGATCCATGATGTCTAAACTGGGAGTCTTGTTGACCATCTGTGTGCTTCTGTTTCCCC  
TTACTGCTGTTCCGCTGGATGGAGATCAACCTGCAGACCAACCTGCAGAGCGTACG  
CAGAACGAGCAGCATCCCTTGTATGATCAGAAAAGAAAGTGTTGCCGGCCGCCATG  
40 CGCCATGAGCTGCGGCATGGCTAGGTGTTGCTATTAATGATAACGTGTTGATGACCA  
ACTTCTCGAG (SEQ ID NO:215)

**Translation:**

45 GSMMSKLGVLLTICVLLFPLTAVPLDGDQPADQPAERTQNEQHPLYDQKRKCCRPPCA  
MSCGMARCCY (SEQ ID NO:216)

**Toxin Sequence:**

Lys-Cys-Cys-Arg-Xaa3-Xaa3-Cys-Ala-Met-Ser-Cys-Gly-Met-Ala-Arg-Cys-Cys-Xaa5-^ (SEQ ID NO:217)

5   **Name:**       Circling  
     **Species:**   textile  
     **Isolated:**   Yes  
     **Cloned:**     Yes

10   **DNA Sequence:**

GAGTCAACCCACTGTCACGCCAAGAGCGGACGCCACAGCTAAGGCAAGAAGGATC  
GATAGCAGTTCATGATGTCTAAACTGGGAGCCTTGTTGACCATCTGTCTACTTCTGT  
TTTCCCTTACTGCTGTTCCGCTGGATGGAGATCAACATGCAGACCAACCTGCACAGC  
GTCTGCAGGACCGCATTCCAAGTGAAGATCATCCCTTATTTGATCCCAACAAACGGT  
15   GTTGCCCGCCGGTGGCATGCAACATGGGATGCAAGCCTTGTTGTGGATGACCAGCTT  
TGTATCGCGGTCTCATGAAGTGTCTAATGAATAAGTAAAACGATTGCAGTTTCGTT  
CAGATTTGCTGTTGTATTTTGGTCTAAAGATTAATGACCAAACCTGTTCTTTTGATCCG  
GATTTTCACGTATTTCTCGATTCCCTATTCAACACTAGATAAGTTAATCACGACAGAT  
CTGATTTTCCATCAATGCCTTGCTTTTGGTCTGTCATATAAATCTTGTTTATATTTAA  
20   TTTCTCGTCACTTTCAACACGCACACACACACACACACACGCGCGCGC (SEQ ID  
NO:218)

**Translation:**

MMSKLGALLTICLLLFSLTAVPLDGDQHADQPAQRLQDRIPTEHPLFDPNKRCCPPVA  
25   CNMGCKPCCG (SEQ ID NO:219).

**Toxin Sequence:**

Cys-Cys-Xaa3-Xaa3-Val-Ala-Cys-Asn-Met-Gly-Cys-Lys-Xaa3-Cys-Cys-Gly-^ (SEQ ID  
NO:220)

30

**Name:**       Scratcher I  
**Species:**   textile  
**Cloned:**     Yes

35

**DNA Sequence:**

GGATCCAGACGACAAAGAAGAGTCAACCCACTGCCACGTCAAGAGCAGAGCCCAC  
AGCTAAGACAAGAAGGATCGATAGCAGTTCATGATGTTTAAACTGGGAGTCTTGTT  
GACCATCTGTCTCCTTCTGTTTCCCTTAATGCTGTTCCGTTGGATGGAGATCAACCT  
40   GCAGACCAACCTGCAGAGCGTCTGCTGGACGACATTTCAATTTGAAAATAATCCCTTT  
TATGATCCCGCCAAACGGTGTGTCAGGACTTGCTTCGGTTGCACACCTTGTTGTGGA  
TGACCAGCCTCATCAAGTGTCTAACGAATAAGTAAAGCGATTGCAGTCTCGTTTCAG  
ATTTACTTTTGTATTCTGGTCTAAAGATTAATGACCAAACCTCTTCTTTTGATCCGGAT  
GTACATATATTTCTCGATTCCCTATCCAACGCTAGATAAGCTAATCACGACAGATCTG  
45   ATTTTCTGTCAATGCCTTGCTTTTGGTCTCTCATATCACTCTTGTTTATATTTAATTT  
CTCGTCACTATATATATATATACACACACACACACACGGAATTCGATTGTCCAGTA  
CCGTTCTTGGGATCGAGGTATTGCTGCGATGGCTTATTCTGTACTCTTTTCTTCTGCG

CTTGATAGTGATGTCTTCTACTCCCATCTGTGCTACCCCTGGCTTGATCTTTGATAGG  
CGTGTGCCCTTCACTGGTTATAAACCCCTCTGATCCTACTCTCTGGACGCCTCGGGG  
GCCCAACCTCCAAATAAAGCGACATCCAATGAAAAA (SEQ ID NO:221)

5 **Translation:**

MMFKLGVLLTICLLLFSLNAVPLDGDQPADQPAERLLDDISFENNPFYDPAKRCCRTCF  
GCTPCCG (SEQ ID NO:222)

**Toxin Sequence:**

10 Cys-Cys-Arg-Thr-Cys-Phe-Gly-Cys-Thr-Xaa3-Cys-Cys-# (SEQ ID NO:223)

**Name:** Tx3.1

**Species:** textile

15 **Cloned:** Yes

**DNA Sequence:**

GGAACAGTCAACCCACAGCCACGCCAAGAGCAGACAGCCACAGCTACGTGAAGA  
AGGGTGGAGAGAGGTTTCATGATGTTGAAAATGGGAGTGGTGCTATTCATCTTTCTGG  
20 TACTGTTTCCCCTGGCAACGCTCCAGCTGGATGCAGATCAACCTGTAGAACGATATG  
CGGAGAACAAACAGCTCCTCAACCCAGATGAAAGGAGGGAAATCCTATTGCCTGCT  
CTGAGGAAGTTCTGCTGTGATTGCAATTGGTGCCACATTTCGGATTGTGAGTGCTGC  
TACGGTTAGCGCCGAACATCCATGGCACTGTGCTGGGCGGTTTCATCCCAACAACG  
ACAGCGTTTGTTGATTTTCATGTATCATTGCGCCACGTCTCTTGTCTAAGAATGACG  
25 AACATGATTGCACTCTGGTTCAGATTTCGTGTTCTTTTCTGACAATAAATGACAAAAC  
TCC (SEQ ID NO:224)

**Translation:**

MMLKMGVVLFIPLVLFPLATLQLDADQPVERYAENKQLLNPDERREILLPALRKFCDS  
30 NWCHISDCECCYG (SEQ ID NO:225)

**Toxin Sequence:**

Phe-Cys-Cys-Asp-Ser-Asn-Xaa4-Cys-His-Ile-Ser-Asp-Cys-Xaa1-Cys-Cys-Xaa5-# (SEQ ID  
NO:226)  
35

**Name:** U031

**Species:** textile

**Isolated:** Yes

40 **Cloned:** Yes

**DNA Sequence:**

CAAGGAACAGTCAACCCACAGCCACGCCAAGAGCAGACAGCCACAGCTACGTGA  
AGAAGGGTGGAGAGAGGTTTCGTGATGTTGAAAATGGGAGTGGTGCTATTCATCTTC  
45 CTGGTACTGTTTCCCCTGGCAACGCTCCAGCTGGATGCAGATCAACCTGTAGAACGA  
TATGCGGAGAACAAACAGCTCCTCAGCCAGATGAAAGGAGGGAAATCATATTGCA  
TGCTCTGGGGACGCGATGCTGTTCTTGGGATGTGTGCGACCAACCGAGTTGTA

CTGCGGTTAGCGCCGAACATCCATGGCGCTGTGCTGGGCGGTTTTATCCCAACAACG  
ACAGCGTTTGTGATTTTCATGTATCATTGCGCCACGTCTCTTGTCTAAGAATGACG  
AACATGATTGCACTCTGGTTCAGATTTTCGTGTTCTTTTCTGACAATAAATGACAAAA  
CNCC (SEQ ID NO:227)

5

**Translation:**

MLKMGVVLFIFLVLFPLATLQLDADQPVERYAENKQLLSPDERREIILHALGTRCCSWD  
VCDHPSCTCCG (SEQ ID NO:228)

10 **Toxin Sequence:**

Cys-Cys-Ser-Xaa4-Asp-Val-Cys-Asp-His-Xaa3-Ser-Cys-Thr-Cys-Cys-# (SEQ ID NO:229)

**Name:** U032

15 **Species:** textile**Isolated:** Yes**Cloned:** Yes**DNA Sequence:**

20 GGATCCATGATGTCTAAACTGGGAGTCTTGTTGACCATCTGTCTGCTTCTGTTTCCCC  
TTACTGCTCTTCCGCTGGATGGAGATCAACCCGACACCAAGCTGCAGAGCGTATG  
CAGGCCGAGCAGCATCCCTTGTTGATCAGAAAAGACGGTGCTGCAAGTTTCCATG  
CCCCGATAGTTGCAGATATTTGTGTTGCGGGTGATGATAACGTGTTGATGACCAACT  
TTCTCGAG (SEQ ID NO:230)

25

**Translation:**

GSMMSKLGVLITICLLFLPTALPLDGDQPADQAAERMQAQHPFLDQKRRCKFP  
DSCRYLCCG (SEQ ID NO:231)

30 **Toxin Sequence:**

Arg-Cys-Cys-Lys-Phe-Xaa3-Cys-Xaa3-Asp-Ser-Cys-Arg-Xaa5-Leu-Cys-Cys-# (SEQ ID NO:232)

35 **Name:** T3.1**Species:** tulipa**Cloned:** Yes**DNA Sequence:**

40 CGACCTCAAGAGGGATCGATAGCAGTTCATGTCTAAACTGGGAGTCTTGTTGACAA  
TCTGTCTGCTTCTGTTTCCCCTTACTGCTCTGCCGATGGATGGAGATGAACCTGCAG  
ACCGACCTGCAGAGCGTATGCAGGACAACATTTTCATCTGAGCAGCATCCCTTGTTTG  
AGGAGAGACACGGATGTTGCAAGGGGCCCCGAAGGATGCTCCTCCAGAGAATGCAG  
ACCCCAACATTGTTGCGGTCGACGATAACGTGTTGAGGGCCAACTTTGTTATCACGG  
45 CTACGTCAAGTGTTTAGTGAATAAGTAAATGATTGCAG (SEQ ID NO:233)

**Translation:**

MSKLGVLLTICLLLFPLTALPMDGDEPADRPAERMQDNISSEQHPLFEERHGCCCKGPEG  
CSSRECRPQHCCGRR (SEQ ID NO:234)

**Toxin Sequence:**

- 5 His-Gly-Cys-Cys-Lys-Gly-Xaa3-Xaa1-Gly-Cys-Ser-Ser-Arg-Xaa1-Cys-Arg-Xaa3-Gln-His-  
Cys-Cys-# (SEQ ID NO:235)

Name: Fi3.1  
10 Species: figulinus  
Cloned: Yes

**DNA Sequence:**

- CAAGAAGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGCTGACCATCT  
15 GTCTGCTTCTGATTCCCCTTACTGCTCTTTCGCTGGATGGAGATCAACCTGCAGACC  
GACCTGCAGAGCGTATGCAGGATGGAATTCATCTGAACAGCATCCCATGTTTGATC  
CCGTACAGACGGTGTTGCCCGTGGCCATGCAACATAGGATGCGTACCTTGTTGTTGAT  
GACCAGTTTTGTTATCGCGGCCTCATCAAATGTCTAATGAATAAGTAAAACGATTGC  
AGT (SEQ ID NO:236)

**Translation:**

MMSKLGVLLTICLLLIPLTALS LDGDQPADRPAERMQDGISSEQHPMFDPVRRCCPWPC  
NIGCVPC (SEQ ID NO:237)

**Toxin Sequence:**

- 25 Cys-Cys-Xaa3-Xaa4-Xaa3-Cys-Asn-Ile-Gly-Cys-Val-Xaa3-Cys-Cys-^ (SEQ ID NO:238)

Name: Fi3.2  
30 Species: figulinus  
Cloned: Yes

**DNA Sequence:**

- CAAGAGGGATCGATAGCAGTTCATGATGTTTAAACTGGGAGTCCTGTTGACCATCTG  
35 TATGCTTCTGTTTCCCTTTACTGCTCTTCCGCTGGATGGAGAGCAACCTGCAGACCA  
ACCTGCAGAGCGCATGCAGTATGACATGTTACGTGCAATGAATCCCTGGTTTGATCC  
CGTCAAAAGGTGCTGCTCGAAGAACTGCGCAGTATGCATCCCTTGTTGCCCGTAACT  
GACCAGCTTGATTATCGCGGCCAAGGCTCTAATGAATAAGTAAAACGATTGCAGT  
(SEQ ID NO:239)

**Translation:**

MMFKLGVLLTICMLLFPTALPLDGEQPADQPAERMQYDMLRAMNPWFDPVKRCCSK  
NCAVCIPCC (SEQ ID NO:240)

**Toxin Sequence:**

- 45 Cys-Cys-Ser-Lys-Asn-Cys-Ala-Val-Cys-Ile-Xaa3-Cys-Cys-Xaa3-^ (SEQ ID NO:241)

**Name:** Fi3.3  
**Species:** *figulinus*  
**Cloned:** Yes

**DNA Sequence:**

CAAGAGGGATCGATAGCAGTTCATGATGTCTAAACTGAGAGTCTTGTTGACCTTATG  
TCTGCTTCTGTTTCCCTTACTGCTCTTCCGCTGAATGAAGATCAACCTGCAGAGCGT  
10 ATGCAGGACGACAATTCATCTGAGCAGCACCCCTTGTATGACCACAAACGAAAGTG  
TTGCCGGTGGCCATGCCCCGCAAGATGCGGCTCTTGTTGCCTGTAATAACGTGTTGG  
CCAACCTTTGTTATCACGGCCACGTCAAATGTTAATGAATAAGTAAAACGATTGCAG  
T (SEQ ID NO:242)

**Translation:**

MMSKLRVLLTLCLLLFPLTALPLNEDQPAERMQDDNSSEQHPLYDHKRKCCRWPCCPAR  
CGSCCL (SEQ ID NO:243)

**Toxin Sequence:**

20 Cys-Cys-Arg-Xaa4-Xaa3-Cys-Xaa3-Ala-Arg-Cys-Gly-Ser-Cys-Cys-Leu-^ (SEQ ID NO:244)

**Name:** Fi3.4  
**Species:** *figulinus*  
**Cloned:** Yes

**DNA Sequence:**

CAAGAGGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGTTGACCTTATG  
TCTGCTTCTGTTTCCCTGACTGCTCTTCCGCTGGATGAAGATCAAGCTGCAGACCG  
30 ACCTGCAGAGCGTATGCAGGGCATGTCATCTGAACAGCATCCCTTCTTTGATCCCGT  
CAAACGGTGTGCGAGTTGTCACGCTGCCTTGGATGCGTCCCTTGTTGCACATCTTA  
ATAACGTGTGGATGACCAACTGTGTTATCACGGCCACGTCAAGTGTCTAATGAATA  
AGTAAAATGATTGCAGT (SEQ ID NO:245)

**Translation:**

MMSKLGVLLTLCLLLFPLTALPLDEDQAADRPAERMQGMSSSEQHPFFDPVKRCCELSR  
CLGCVPCCTS (SEQ ID NO:246)

**Toxin Sequence:**

40 Cys-Cys-Xaa1-Leu-Ser-Arg-Cys-Leu-Gly-Cys-Val-Xaa3-Cys-Cys-Thr-Ser-^ (SEQ ID NO:247)

**Name:** Fi3.5  
**Species:** *figulinus*  
**Cloned:** Yes

**DNA Sequence:**

62

CAAGAGGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGTTGACCTTATG  
TCTGCTTCTGTTTCCCCTGACTGCTCTCCGCTGGATGAAGATCAACCTGCAGACCG  
ACCTGCAGAGCGTATGCAGGGCATGTCATCTGAACAGCATCCCTTCTTTGATCCCGT  
CAAACGGTGTTGCGAGTTGTCAAAATGCCATGGATGCGTCCCTTGTTGCATACCTTA  
5 ATAACGTGCGGATGACCAACTGTGTTATCACGGCCACGTCAAGTGTCTAATGAATA  
AGTAAATGATTGCAGT (SEQ ID NO:248)

**Translation:**

MMSKLGVLTLCLLLFPLTALPLDEDQPADRPAERMQMSSEQHPFFDPVKRCCELSK  
10 CHGCVPCIP (SEQ ID NO:249)

**Toxin Sequence:**

Cys-Cys-Xaa1-Leu-Ser-Lys-Cys-His-Gly-Cys-Val-Xaa3-Cys-Cys-Ile-Xaa3-^ (SEQ ID NO:250)

15

**Name:** Qc3.2  
**Species:** quercinus  
**Cloned:** Yes

**DNA Sequence:**

CAAGAGGGATCGATAGCAGTTCATGATGTCTAAACTCGGAGTCTTGTTGACCATCTG  
TCTGGTTCTGTTTCCCCTTACAGCTCTTCAGCTGGATGGAGATCAACCTGCAGACCG  
ACCTGCAGAGCGTACGCAGGACATTTTCATCTGAACAGTATCGAAAGTTTGATCAGA  
GACAGAGGTGTTGCCGGTGGCCATGCCCCGGTAGTTGCAGATGCTGCCGTTATCGTT  
25 AACGTGTTGGTGACCAGCTTTGTTATCACGACCACGCCAAGTGTCTAACGAATAAGT  
AAAATGATTGCAGT (SEQ ID NO:251)

**Translation:**

MMSKLGVLTLICLVLFPLTALQLDGDQPADRPAERTQDISSEQYRKFDQRQRCCRWP  
30 GSCRCCRYR (SEQ ID NO:252)

**Toxin Sequence:**

Xaa2-Arg-Cys-Cys-Arg-Xaa4-Xaa3-Cys-Xaa3-Gly-Ser-Cys-Arg-Cys-Arg-Xaa5-Arg-^  
(SEQ ID NO:253)

35

**Name:** Qc3.3  
**Species:** quercinus  
**Cloned:** Yes

40

**DNA Sequence:**

CAAGAGGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGTTGACCATCTG  
TCTGCTTCTGTTTCCCCTTACTGCTCTTCCACTGGATGGAGATCAACCTGCAGATCAA  
TCTGCAGAGCGACCTGCAGAGCGTACGCAGGACGACATTCAGCAGCATCCGTTATA  
45 TGATCCGAAAAGAAGGTGTTGCCGTTATCCATGCCCCGACAGCTGCCACGGATCTTG  
CTGCTATAAGTGATAACATGTTGATGGCCAGCTTTGTTATCACGGCCACGTCAAGTG  
TCTAATGAATAAGTAAACGATTGCAGT (SEQ ID NO:254)

**Translation:**

MMSKLGVLLTICLLLFPLTALPLDGDQPADQSAERPAERTQDDIQHPLYDPKRRCRY  
PCPDSCHGSCCYK (SEQ ID NO:255)

5

**Toxin Sequence:**

Arg-Cys-Cys-Arg-Xaa5-Xaa3-Cys-Xaa3-Asp-Ser-Cys-His-Gly-Ser-Cys-Cys-Xaa5-Lys-^ (SEQ  
ID NO:256)

10

**Name:** Wi3.1  
**Species:** wittigi  
**Cloned:** Yes

15 **DNA Sequence:**

GGATCCATGATGTCTAAACTGGGAGTCTTGTGACCATCTGTCTGCTTCTGTTTCCCA  
TTACTGCTCTTCCGGTGGGTGGAGATCAGCCTGCAGACCGACTTGCAGAGCGTATGC  
AGGACGACACTTCATCTGAGCAGCATCCCTTTGAAAAGAGACTACCATCATGTTGC  
GACTTTGAGAGGCTTTGCGTAGTACCAGCATGCATACGTCATCAGTGTTGCACAGGA  
20 TAACGTGTTGATGACCAACTTTCTCGAG (SEQ ID NO:257)

**Translation:**

MMSKLGVLLTICLLLFITALPVGGDQPADRLAERMQDDTSSEQHPFEKRLPSCCDFERL  
CVVPACIRHQCCTG (SEQ ID NO:258)

25

**Toxin Sequence:**

Leu-Xaa3-Ser-Cys-Cys-Asp-Phe-Xaa1-Arg-Leu-Cys-Val-Val-Xaa3-Ala-Cys-Ile-Arg-His-Gln-  
Cys-Cys-Thr-# (SEQ ID NO:259)

30

**Name:** bt3a  
**Species:** betulinus  
**Isolated:** Yes

35 **Toxin Sequence:**

Cys-Cys-Lys-Gln-Ser-Cys-Thr-Thr-Cys-Met-Xaa3-Cys-Cys-Xaa4-^ (SEQ ID NO:260)

40

**Name:** T3.2  
**Species:** tulipa  
**Cloned:** Yes

**DNA Sequence:**

45 GGATCCATGATGTCTAAACTGGGAGTCTTGTGACAACTCTGTCTGCTTCTGTTTCCCC  
TTACTGCTCTGCCGATGGATGGAGATGAACCTGCAGACCGACCTGCAGAGCGTATG  
CAGGACAACATTTTCATCTGAGCAGCATCCCTTGTGTTGAGGAGAGACACGGATGTTG



64

CGAGGGGCGGAAGGGATGCTCCTCCAGAGAATGCAGACCCCAACATTGTTGCGGTC  
GACGATAACGTGTTGATGACCAACTNTCTCGAG (SEQ ID NO:261)

**Translation:**

5 MMSKLGVLITICLLLFPLTALPMDGDEPADRPAERMQDNISSEQHPLFEERHGCCEGPK  
GCSSRECRPQHCCGRR (SEQ ID NO:262)

**Toxin Sequence:**

10 His-Gly-Cys-Cys-Xaa1-Gly-Xaa3-Lys-Gly-Cys-Ser-Ser-Arg-Xaa1-Cys-Arg-Xaa3-Gln-His-  
Cys-Cys-# (SEQ ID NO:263)

Name: A3.5  
Species: aurisiacus  
15 Cloned: Yes

**DNA Sequence:**

GGATCCATGATGTCTAAACTGGGAGTCTTGTTGACCATCTGTCTACTTCTGTTTCCCC  
TTACTGCTTTTCCGATGGATGGAGATCAACCTGCAGACCAACCTGCAGATCGTATGC  
20 AGGACGACATTTTCATCTGAGCAGTATCCCTTGTTTGATAAGAGACAAAAGTGTGCA  
CTGGGAGGAAGGGGTCATGCTCCGGCAAAGCATGCAAAAATCTCAAATGTTGCTCT  
GGACGATAACGTGTTGATGACCAACTTTCTCGAN (SEQ ID NO:264)

**Translation:**

25 MMSKLGVLITICLLLFPLTAFPMGDGQPADQPADRMQDDISSEYPLFDKRQKCCTGR  
KGSCSGKACKNLKCCSGR (SEQ ID NO:265)

**Toxin Sequence:**

30 Xaa2-Lys-Cys-Cys-Thr-Gly-Arg-Lys-Gly-Ser-Cys-Ser-Gly-Lys-Ala-Cys-Lys-Asn-Leu-Lys-  
Cys-Cys-Ser-# (SEQ ID NO:266)

Name: Bt3.5  
Species: betulinus  
35 Cloned: Yes

**DNA Sequence:**

GGATCCATGATGTCTAAACTGGGAGTCTTGTTGACCATCTGTCTGCTTCTGTTTCCCC  
TTACTGCTGTTCCGTTGGATGGAGATCAACCTGCAGACCAACCTGCAGAGCGTATGC  
40 AGAACGAGCAGCATCCCTCGTTTGATCAGAAAAGAAGGTGCTGCCGGTGGCCATGC  
CCCAGTATATGCGGCATGGCTAGGTGTTGCTTCGTCATGATAACGTGTTGATGACCA  
ACTTTCTCGAG (SEQ ID NO:267)

**Translation:**

45 MMSKLGVLITICLLLFPLTAVPLDGDQPADQPAERMQNEQHPSFDQKRRCRWPCCPSIC  
GMARCCFVMITC (SEQ ID NO:268)

**Toxin Sequence:**

Arg-Cys-Cys-Arg-Xaa4-Xaa3-Cys-Xaa3-Ser-Ile-Cys-Gly-Met-Ala-Arg-Cys-Cys-Phe-Val-Met-Ile-Thr-Cys-^ (SEQ ID NO:269)

5

**Name:** Bt3.6  
**Species:** betulinus  
**Cloned:** Yes

10

**DNA Sequence:**

GGATCCATGATGTCTAAACTGGGAGTCTTGTTGATCATCTGTCTGCTTCTGTTTCCCC  
TTACTGCTGTTCCGCTGGATGGAGATCAGCCTGCAGAGCGTACGCAGATCGAGCAG  
CATCCCTTGTTTGACCAGAAAAGAAGGTGTTGCCGGTGGCCATGCCCCAGTAGATG  
CGGCATGGCTAGGTGTTGCTTCGTCATGATAACGTGTTGATGANCGACCTCTCNAG  
(SEQ ID NO:270)

15

**Translation:**

MMSKLGVLLIICLLLFPLTAVPLDGDQPAERTQIEQHPLFDQKRRCCRWP CPSRCGMAR  
CCFVMITC (SEQ ID NO:271)

20

**Toxin Sequence:**

Arg-Cys-Cys-Arg-Xaa4-Xaa3-Cys-Xaa3-Ser-Arg-Cys-Gly-Met-Ala-Arg-Cys-Cys-Phe-Val-Met-Ile-Thr-Cys-^ (SEQ ID NO:272)

25

**Name:** Pr3.1  
**Species:** parius  
**Cloned:** Yes

30

**DNA Sequence:**

GGATCCATGATGTCTAAACTGGGAGTCTTGTTGACCATCTGTCTGCTTCTGTTTCCCC  
TTACTGCTCTTCCGATGGATGGTGATCAACCTGCAGACCGACTTG TAGAGCGTATGC  
AGGACAACATTTTCATCTGAGCAGCATCCCTTCTTTGAAAAGAGAAGAGGAGGCTGT  
TGCACACCTCCGAAGAAATGCAAAGACCGAGCCTGCAAACCTGCACGTTGCTGCGG  
CCCAGGATAACGTGTTGATGACCAACTTTCTCGCC (SEQ ID NO:273)

35

**Translation:**

MMSKLGVLLTICLLLFPLTALPMDGDQPADRLVERMQDNISSEQHPFFEKRRGGCCTPP  
KKCKDRACKPARCCGPG (SEQ ID NO:274)

40

**Toxin Sequence:**

Arg-Gly-Gly-Cys-Cys-Thr-Xaa3-Xaa3-Lys-Lys-Cys-Lys-Asp-Arg-Ala-Cys-Lys-Xaa3-Ala-Arg-Cys-Cys-Gly-Xaa3-# (SEQ ID NO:275)

45

**Name:** Pr3.2  
**Species:** parius  
**Cloned:** Yes

5 **DNA Sequence:**

GGATCCATGATGTCTAAACTGGGAGTCTTGTTGACCATCTGTCTGCTTCTGTTTCCCC  
TTACTGCTCTTCCGATGGATGGTGATCAACCTGCAGACCGACTTGTAAGAGCGTATGC  
AGGACAACATTTTCATCTGAGCAGCATCCCTTCTTTGAAAAGAGAAGAGGCTGTTGC  
ACACCTCCGAGGAAATGCAAAGACCGAGCCTGCAAACCTGCACGTTGTTGCGGCCC  
10 AGGATAACGTGTTGATGACCAACTTTCTCGAG (SEQ ID NO:276)

**Translation:**

MMSKLGVLTTICLLLFPLTALPMDGDQPADRLVERMQDNISSEQHPFFEKRRGCCTPPR  
KCKDRACKPARCCGPG (SEQ ID NO:277)

15

**Toxin Sequence:**

Arg-Gly-Cys-Cys-Thr-Xaa3-Xaa3-Arg-Lys-Cys-Lys-Asp-Arg-Ala-Cys-Lys-Xaa3-Ala-Arg-  
Cys-Cys-Gly-Xaa3-# (SEQ ID NO:278)

20

**Name:** Ct3.1  
**Species:** coronatus  
**Cloned:** Yes

25 **DNA Sequence:**

GGATCCATGATGTCTAAACTGGGAGTCTTGTTGACCATCTGTCTGCTTCTGTTTCCAA  
TTACTGCCCTTCCGCTGGATGAAGATCAACCTGCAGACCGACCTGCAGAGCGTATGC  
AGGACATTGCAACTGAACAGCATCCCTTGTTTGATCCCGTCAAACGGTGCTGCGATT  
GGCCATGCATCCCAGGATGCACCCCTTGTTGCTTGCCTTGATAACGTGTTGATGACC  
30 AACTTTCTCGAG (SEQ ID NO:279)

**Translation:**

MMSKLGVLTTICLLLFPTALPLDEDQPADRPAERMQDIATEQHPLFDPVKRCCDWPCIP  
GCTPCCLP (SEQ ID NO:280)

35

**Toxin Sequence:**

Cys-Cys-Asp-Xaa4-Xaa3-Cys-Ile-Xaa3-Gly-Cys-Thr-Xaa3-Cys-Cys-Leu-Xaa3-^ (SEQ ID  
NO:281)

40

**Name:** Ms3.1  
**Species:** musicus  
**Cloned:** Yes

45 **DNA Sequence:**

GGATCCATGATGTCTAAACTGGGAGTCCTGTTGACCATCTGTCTGCTTCTGTTTCCTC  
TTTCTGCTCTTCCGATGGATGAAGATCAACTTGCAGACCTACCTGCAGAGCGTATGC

GGGACACTGCAACTGTAGATCATCCCTCCTATGATCCTGACAAAGCGTGCTGCGAG  
CAGAGCTGTACAACATGCTTTCCGTGCTGCTAGCCTTGAACACAGTAACGTGTTGAT  
GACCAACTTTCTCGAG (SEQ ID NO:282)

5 **Translation:**

MMSKLGVLITICLLLFPLSALPMDEDQLADLPAERMRTATVDHPSYDPDKACCEQSC  
TTCFPCC (SEQ ID NO:283)

**Toxin Sequence:**

10 Ala-Cys-Cys-Xaa1-Gln-Ser-Cys-Thr-Thr-Cys-Phe-Xaa3-Cys-Cys-^ (SEQ ID NO:284)

**Name:** bt3b

**Species:** betulinus

15 **Isolated:** Yes

**Toxin Sequence:**

20 Ala-Cys-Cys-Xaa1-Gln-Ser-Cys-Thr-Thr-Cys-Met-Xaa3-Cys-Cys-^ (SEQ ID NO:285)

**Name:** bt3c

**Species:** betulinus

25 **Isolated:** Yes

**Toxin Sequence:**

Cys-Cys-Xaa1-Gln-Ser-Cys-Thr-Thr-Cys-Met-Xaa3-Cys-Cys-Xaa4-? (SEQ ID NO:286)

30 **Name:** Pn3.2

**Species:** pennaceus

**Cloned:** Yes

**DNA Sequence:**

35 GGATCCATGATGTCTAAACTGGGAGTCTTGTTGACCATCTGTCTGCTTCTGTTTCCCC  
TTACTGCTCTTCCGCTGGATGGAGATCAACCTGCATACCAAGCTGCAGAGCGTATGC  
AGGCCGAGCATCATCCCTTGTTTGATCAGAAAAGACGGTGCTGCAAGTTTCCATGCC  
CCGATAGTTGCAAATATTTGTGTTGCGGGTGATGATAACATGTTGATGACCAACTTT  
CTTGAG (SEQ ID NO:287)

40

**Translation:**

MMSKLGVLITICLLLFPLTALPLDGDQPAYQAAERMQAEBHPLFDQKRRCKFPDPS  
CKYLCCG (SEQ ID NO:288)

45 **Toxin Sequence:**

Arg-Cys-Cys-Lys-Phe-Xaa3-Cys-Xaa3-Asp-Ser-Cys-Lys-Xaa5-Leu-Cys-Cys-# (SEQ ID NO:289)

Name: Pu3.2  
Species: pulicarius  
Cloned: Yes

**DNA Sequence:**

GGATCCATGATGTCTAAACTGGGAGTCTTGTTGACCATCTGTCTGCTTCTGTTTCCCC  
TTACTGCTCTTCCGATGGATGGTGATCAACTTGCAGACCGACTTGTAGAGCGTATGC  
10 AGGACAACATTTTCATCTGAGCAGCATCCCTTCTTTGATCCCGTCAAACGGTGTGCG  
TCAGCTGTTACATGGGATGCATCCCTTGTGCTTCTAGTAATAACGTGTTGATGACC  
AACTTTCTCGAG (SEQ ID NO:290)

**Translation:**

15 MMSKLGVLITICLLLFPLTALPMDGDQLADRLVERMQDNISSEQHPFFDPVKRCCVSCY  
MGCIPCCF (SEQ ID NO:291)

**Toxin Sequence:**

20 Cys-Cys-Val-Ser-Cys-Xaa5-Met-Gly-Cys-Ile-Xaa3-Cys-Cys-Phe-^ (SEQ ID NO:292)

Name: Pu3.3  
Species: pulicarius  
Cloned: Yes

**DNA Sequence:**

GGATCCATGATGTCTAAACTGGGAGTCTTGTTGACCGTCTGTCTGCTTCTGTGTCCCC  
TTACTGCTCTTCCACTGGATGAAGATCAACTTGCAGACCGACCTGCAGAGCGTATGC  
AGGATGACACTTCAGCTGCACAGATTTTCGGGTTTGATCCCGTCAAACGGTGTGCTGCA  
30 AATTGCTATGCTACTCGGGATGCACTCCTTGTGCCATATTTGATAACGTGTTGATG  
ACCAACTTTCTCGAG (SEQ ID NO:293)

**Translation:**

35 MMSKLGVLITVCLLLCPLTALPLDEDQLADRPAERMQDDTSAAQIFGFDPVKRCCKLL  
CYSGCTPCCHI (SEQ ID NO:294)

**Toxin Sequence:**

Cys-Cys-Lys-Leu-Leu-Cys-Xaa5-Ser-Gly-Cys-Thr-Xaa3-Cys-Cys-His-Ile-^ (SEQ ID NO:295)

40 Name: Ra3.2  
Species: rattus  
Cloned: Yes

**DNA Sequence:**

45 GGATCCATGATGTCTAAACTGGGAGTCTTGTTGACCATCTGTCTGCTTGTGTTTCCGC  
TTACTGCTCTTCCGATGGATGGTGATCAACCTGCAGACCGACTTGTAGAGCGTATAC

AGGACAACATTTTCATCTGAGCAGCATCCCTTCTTTGAAAAGAGAAGAGGCTGTTGC  
GCACCTCCGAGGAAATGCAAAGACCGAGCCTGCAAACCTGCACGTTGCTGCGGCCC  
AGGATAACGTGTTGATGACCAACTTTCTCGAG (SEQ ID NO:296)

5 **Translation:**

MMSKLGVLITICLLVFPLTALPMDGDQPADRLVERIQDNISSEQHPFFEKRRGCCAPPRK  
CKDRACKPARCCGPG (SEQ ID NO:297)

**Toxin Sequence:**

10 Arg-Gly-Cys-Cys-Ala-Xaa3-Xaa3-Arg-Lys-Cys-Lys-Asp-Arg-Ala-Cys-Lys-Xaa3-Ala-Arg-  
Cys-Cys-Gly-Xaa3-# (SEQ ID NO:298)

**Name:** Sm3.3

15 **Species:** stercusmuscarum

**Cloned:** Yes

**DNA Sequence:**

GGATCCATGATGTCTAAACTGGGAGTCTTGTTGACAATCTGTCTGCTTCTGTTTCCCC  
20 TTATTGCTCTTCCGCTGGATGGAGATCAACCTGCAGACCGACCTGCAGAGCGTATGC  
AGGACGACATTTTCATCTGAGAAGCATCCCTTGTTTGATAAGAGACAACGGTGTTC  
AATGGGCGGAGGGGATGCTCCAGCAGATGGTGCAGAGATCACTCACGTTGTTGCGG  
TCGACGATAACGTGTTGATGACCAACTTTCTCGAG (SEQ ID NO:299)

25 **Translation:**

MMSKLGVLITICLLLPLIALPLDGDQPADRPAERMQDDISSEKHPLFDKRQRCCNGRR  
GCSSRWCRDHSRCCGRR (SEQ ID NO:300)

**Toxin Sequence:**

30 Xaa2-Arg-Cys-Cys-Asn-Gly-Arg-Arg-Gly-Cys-Ser-Ser-Arg-Xaa4-Cys-Arg-Asp-His-Ser-Arg-  
Cys-Cys-# (SEQ ID NO:301)

**Name:** Eb3.1

35 **Species:** ebraeus

**Cloned:** Yes

**DNA Sequence:**

GGATCCATGATGTCTAAACTGGGAGTCTTGTTGACCATCTGTCTGCTTCTGTTTCCCC  
40 TTACTGCTCTTCCACTGGATGAAGGTCAACCTGCAGACCTACCTGCAGAGCGTATGC  
AGGACATTGCAACTGAACAGCATCCCTTGTTTGATCCTGTCAAACGGTGTTCGAGC  
AGCCATGCTACATGGGATGCATCCCTTGTTGCTTCTAATAATAACGTGTTGATGACC  
AACTTTCTCGAG (SEQ ID NO:302)

45 **Translation:**

MMSKLGVLITICLLLPLTALPLDEGQPADLPAERMQDIATEQHPLFDPVKRCCEQPCY  
MGCIPCCF (SEQ ID NO:303)

**Toxin Sequence:**

Cys-Cys-Xaa1-Gln-Xaa3-Cys-Xaa5-Met-Gly-Cys-Ile-Xaa3-Cys-Cys-Phe-^ (SEQ ID NO:304)

5

**Name:** Eb3.2  
**Species:** ebraeus  
**Cloned:** Yes

10 **DNA Sequence:**

GGATCCATGATGTCTAAACTGGGAGTCTTGTTGACCATCTGTCTGCTTCTGTTTCCCC  
TTACTGCTCTTCCACTGGATGAAGATCAACCTGCAGACCTACCTGCAGAGCGTATGC  
AGGACATTGCAACTGAACAGCATCCCTTGTTTGATCCTGTCAAACGGTGCTGCGCGC  
AGCCATGCTACATGGGATGCATCCCTTGTTGCTTCTAATAATAACGTGTTGATGACC  
15 AACTTTCTCGAG (SEQ ID NO:305)

**Translation:**

MMSKLGVLLTICLLLFPLTALPLDEDQPADLPAERMQDIATEQHPLFDPVKRCCAQPCY  
MGCI PCCF (SEQ ID NO:306)

20

**Toxin Sequence:**

Cys-Cys-Ala-Gln-Xaa3-Cys-Xaa5-Met-Gly-Cys-Ile-Xaa3-Cys-Cys-Phe-^ (SEQ ID NO:307)

25

**Name:** Fd3.2  
**Species:** flavidus  
**Cloned:** Yes

**DNA Sequence:**

30 GGATCCATGATGTCTAAACTGGGAGTCTTGTTGACCATCTGTCTGCTTCTGTTTCCCC  
TTACTGCTGTTCCGTTGGATGGAGATCAACCTGCAGACCAGCCTGCAGAGCGTATGC  
AGAACGAGCAGCATCCCTTGTTTGATCAGAAAAGAAGGTGCTGCCGGTGGCCATGC  
CCCAGTATATGCGGCATGGCTAGGTGTTGCTCGTCATGATAACGTGTTGATGACCAA  
CTTTCTCGAG (SEQ ID NO:308)

35

**Translation:**

MMSKLGVLLTICLLLFPLTAVPLDGDQPADQPAERMQNEQHPLFDQKRRCCRWPCPSIC  
GMARCCSS (SEQ ID NO:309)

40

**Toxin Sequence:**

Arg-Cys-Cys-Arg-Xaa4-Xaa3-Cys-Xaa3-Ser-Ile-Cys-Gly-Met-Ala-Arg-Cys-Cys-Ser-Ser-^  
(SEQ ID NO:310)

45

**Name:** Mf3.1  
**Species:** miliaris  
**Cloned:** Yes

**DNA Sequence:**

GGATCCATGATGTCTAAACTGGGAGTCTTGTTGACCATCTGTCTGCTTCTGTTTCCAA  
TTACTGCCCTTCCACTGGATGAAGATCAACCTGCAGACCGACCTGCAGAGCGTATGC  
5 AGGACATTGCAACTGAACAGCATCCCTTGTTTGATCCCGTCAAACGGTGTGCGATT  
GGCCATGCAGCGCAGGATGCTACCCTTGTTGCTTCCCTTAATAACGTGTTGATGACC  
AACTNANGNAAAAAAAA (SEQ ID NO:311)

**Translation:**

10 MMSKLGVLITICLLLPITALPLDEDQPADRPABERMQDIATEQHPLFDPVKRCCDWPCS  
AGCYPCCFP (SEQ ID NO:312)

**Toxin Sequence:**

15 Cys-Cys-Asp-Xaa4-Xaa3-Cys-Ser-Ala-Gly-Cys-Xaa5-Xaa3-Cys-Cys-Phe-Xaa3-^ (SEQ ID  
NO:313)

**Name:** Mf3.2  
**Species:** miliaris  
20 **Cloned:** Yes  
**Notes:**

**DNA Sequence:**

GGATCCATGATGTCTAAACTGGGAGTGGTGCCATTTCGTCTTTCTGGTCCTGTTTCCCC  
25 TGGCAACACTCCAACCTGGATGCAGATCAACCTGCAGACCGACCTGCGCGTAAAAAG  
GGCATTGCAACTAAACGGCATCCCTTGCTGTGATCCTGTCAGAGGGTGTGCCCCTCCA  
ATGTGCACACCATGCTTCCCTTGCTGTTTTCGTTAATAACGTGTTGATGNATGATGN  
AN (SEQ ID NO:314)

**Translation:**

30 MMSKLGVPFVFLVLFPLATLQLDADQPADRPARKKGIATKRHPLSDPVRGCCPPMCTPCFPCC  
FR (SEQ ID NO:315)

**Toxin Sequence:**

35 Gly-Cys-Cys-Xaa3-Xaa3-Met-Cys-Thr-Xaa3-Cys-Phe-Xaa3-Cys-Cys-Phe-Arg-^ (SEQ ID  
NO:316)

**Name:** Af3.1  
40 **Species:** ammiralis  
**Cloned:** Yes

**DNA Sequence:**

CAAGAGGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGTTGACCATCTG  
45 TCTGCTTCTGTTTCCCCTTACTGCTCTTCCGCTGGATGGAGATCAACCTGCAGACCA  
AGCTGCAGAGCGTATGCAGGCCGAGCAGCATCCCTTGTTTGATCAGAAAAGACGGT  
GTTGCAGGTTTCCATGCCCCGATACTTGACAGACATTTGTGTTGCGGGTGATGATAAC



GTGCTGATGACCCACTTTGTCATCACGGCTACGTCAAGTGTCTAATGAATAAGTAAA  
ATGATTGCAGT (SEQ ID NO:317)

**Translation:**

5 MMSKLGVLITICLLLFPLTALPLDGDQPADQAAERMQAEOHPLFDQKRRCCRFPCPDT  
CRHLCCG (SEQ ID NO:318)

**Toxin Sequence:**

10 Arg-Cys-Cys-Arg-Phe-Xaa3-Cys-Xaa3-Asp-Thr-Cys-Arg-His-Leu-Cys-Cys-# (SEQ ID  
NO:319)

**Name:** Af3.2

**Species:** ammimalis

15 **Cloned:** Yes

**DNA Sequence:**

CAAGAGGGATCGATAGCAGTTCATGATGTTTAAACTGGGAGTCTTGCTGACCATCTG  
TCTACTTCTGTTTTCCCTTAATGCTGTTCCGCTGGATGGAGATCAACCTGCAGACCA  
20 ACCTGCAGAGCGTCTGCTGGACGACATTTATCTGAAAATAATCCCTTTTATGATCC  
CGCCAAACGGTGTTCATGACTTGCTTCGGTTGCACACCTTGTGTGGATGACCAGC  
CTCATCAAGTGTCTAACGAATAAGTAAAACGATTGCAGT (SEQ ID NO:320)

**Translation:**

25 MMFKLGVLITICLLLFSLNAVPLDGDQPADQPAERLLDDISSENNPFYDPAKRCCMTCTF  
GCTPCCG (SEQ ID NO:321)

**Toxin Sequence:**

30 Cys-Cys-Met-Thr-Cys-Phe-Gly-Cys-Thr-Xaa3-Cys-Cys-# (SEQ ID NO:322)

**Name:** Af3.3

**Species:** ammimalis

35 **Cloned:** Yes

**DNA Sequence:**

CAAGAAGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGCCTTGTTGACCATCT  
GTCTACTTCTGTTTTCCCTTACTGCTGTTCCGCTGGATGGAGATCAACATGCAGACC  
AACCTGCAGAGCGTCTGCAGGACCGCCTTCCAACCTGAAAATCATCCCTTATATGATC  
40 CCGTCAAACGGTGTTCGATGATTCGGAATGCGACTATTCTTGCTGGCCTTGCTGTA  
TTTTTTCATAACCTTTGTTATCGCGGCCTCATCCTAGTGTCAAATGAATAAGTAAAA  
CGATTGCAGT (SEQ ID NO:323)

**Translation:**

45 MMSKLGALLTICLLLFSLTAVPLDGDQHADQPAERLQDRLPTENHPLYDPVKRCCDDSE  
CDYSCWPCCIFS (SEQ ID NO:324)

**Toxin Sequence:**

Cys-Cys-Asp-Asp-Ser-Xaa1-Cys-Asp-Xaa5-Ser-Cys-Xaa4-Xaa3-Cys-Cys-Ile-Phe-Ser-^ (SEQ ID NO:325)

5

**Name:** Af3.4  
**Species:** ammimalis  
**Cloned:** Yes

10

**DNA Sequence:**

CAAGAGGGATCGATAGCAGTTCATGATGTTTAAACTCGGAGTCTTGCTGACCATCTG  
TCTACTTCTGTTTTCCCTAATGCTGTTCCGCTGGATGGAGATCAACATGCAGACCAA  
CCTGCAGAGCGTCTGCAGGACCGCCTTCCAACCTGAAAATCATCCCTTATATGATCCC  
GTCAAACGGTGTTGCAGGTTGTTATGCCTCAGTTGCAACCCTTGTTGTGGATGACCA  
GCTTTGTTATCACGGCCTCATCAAGTGTCTAATGAATAAGTAAAACGATTGCAGT  
(SEQ ID NO:326)

15

**Translation:**

MMFKLGVLLTICLLLFSLIAVPLDGDQHADQPAERLQDRLPTENHPLYDPVKRCCRLLC  
LSCNPCCG (SEQ ID NO:327)

20

**Toxin Sequence:**

Cys-Cys-Arg-Leu-Leu-Cys-Leu-Ser-Cys-Asn-Xaa3-Cys-Cys-# (SEQ ID NO:328)

25

**Name:** Af3.6  
**Species:** ammimalis  
**Cloned:** Yes

30

**DNA Sequence:**

CAAGAAGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGCCTTGTTGACCATCT  
GTCTACTTCTGTTTTCCCTTACTGCTGTTCCGCTGGATGGAGATCAACATGCAGACC  
AACCTGCAGAGCGTCTGCAGGACCGCATTCCAACCTGAAGATCATCCCTTATTTGATC  
CCAACAAACGGTGTTGCGATGATTCGGAATGCGGCTATTCATGCTGGCCTTGCTGTT  
ATGGATAAGCTTTGTTATCGCGGCCTCATCCAGTGTCAACGAATAAGTAAAACGATT  
GCAGT (SEQ ID NO:329)

35

**Translation:**

MMSKLGALLTICLLLFSLTAVPLDGDQHADQPAERLQDRIPTEHDHPLFDPNKRCCDDSE  
CGYSCWPCCYG (SEQ ID NO:330)

40

**Toxin Sequence:**

Cys-Cys-Asp-Asp-Ser-Xaa1-Cys-Gly-Xaa5-Ser-Cys-Xaa4-Xaa3-Cys-Cys-Xaa5-# (SEQ ID NO:331)

45

**Name:** Sf3.1  
**Species:** spurius  
**Cloned:** Yes

5 **DNA Sequence:**

CAAGAAGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGCTGACCATCT  
GTCTGCTTCTGTTTCCACGTACTTCTCTTCCGCTGGATGGAGATCAACCTGCAGTCCG  
ATCTGCAAAGCGTATGCATTATCTATACAGCGTCGTTTCTTTGATCCCGTCAAACG  
GTGTTGCCCTAGATGCAGCGAGTGCAACCCCTGTTGTGGATGACCAGCTTTGTCATC  
10 GCGGCCTCATTAAGTGTCTAATGAATAAGTAAAATGATTGCAGT (SEQ ID NO:332)

**Translation:**

MMSKLGVLLTICLLLPRTSLPLDGDQPAVRS AKRMHSSIQRFFDPVKRCCPRCSECNP  
CCG (SEQ ID NO:333)

15

**Toxin Sequence:**

Cys-Cys-Xaa3-Arg-Ser-Xaa1-Cys-Asn-Xaa3-Cys-Cys-# (SEQ ID NO:334)

20 **Name:** Om3.1  
**Species:** omaria  
**Cloned:** Yes

**DNA Sequence:**

25 CAAGAGGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTCGTTGACCATCT  
GTCTACTTCTATTTTCCCTTACTGCTGTTCCGCTTGATGGAGATCAACATGCAGACCA  
ACCTGCAGAGCGTCTGCAGGGCGACATTTTATCTGAAAAGCATCCCTTATTTAATCC  
CGTCAAACGGTGTTGCGATGAGGAAGAATGCAGCAGTGCATGCTGGCCTTGTTGTT  
GGGGGTGATCAGCTTTGTTATCGCGGCCTCATCAAGTGTCTAATGAATAAGTAAAAT  
30 GATTGCAGT (SEQ ID NO:335)

**Translation:**

MMSKLGVS LTICLLLFSLTAVPLDGDQHADQPAERLQGDILSEKHPLFNPVKRCCDEEE  
CSSACWPCCWG (SEQ ID NO:336)

35

**Toxin Sequence:**

Cys-Cys-Asp-Xaa1-Xaa1-Xaa1-Cys-Ser-Ser-Ala-Cys-Xaa4-Xaa3-Cys-Cys-Xaa4-# (SEQ ID  
NO:337)

40

**Name:** Om3.2  
**Species:** omaria  
**Cloned:** Yes

45 **DNA Sequence:**

CAAGAAGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGTTGATCATCTG  
TCTACTTCTGTGTCCCTTACTGCTGTTCTGGAGGATGGAGATCAACCTGCAGACCG

ACCTGCAGAGCGTATGCAGGACGACATTTCAACTGAGCATCATCCCTTTTATGATCC  
CGTCAAACGGTGTGCAAGTACGGGTGGACATGCTTGCTAGGATGCACTCCTTGTGA  
TTGTTGACCAGTTTTGTTATCGCGGCCTCGTCAAGTGTCTAATGAATAAGTAAAACG  
ATTGCAGT (SEQ ID NO:338)

5

**Translation:**

MMSKLGVLIIICLLCPLTAVLEDGDQPADRPAERMQDDISTEHHFPFYDPVKRCCKYG  
WTCLLGCTPCDC (SEQ ID NO:339)

10 **Toxin Sequence:**

Cys-Cys-Lys-Xaa5-Gly-Xaa4-Thr-Cys-Leu-Leu-Gly-Cys-Thr-Xaa3-Cys-Asp-Cys-^ (SEQ ID NO:340)

15 **Name:** Om3.3  
**Species:** omaria  
**Cloned:** Yes

**DNA Sequence:**

20 CAAGAGGGATCGATAGCAGTTCATGATGTCTATACTGGGAGTCTTGTTGATCATCTG  
TCTACTTCTGTGTCCCCTTACTGCTGTTCTGGAGGATGGAGATCAACCTGCAGACCG  
ACCTGCAGAGCGTATGCAGGACGGCATTTCATCTGAACATCATCCCTTTTTGGATCC  
CGTCAAACGGTGTGTCATCTATTGGCATGCCGCTTTGGATGCTCGCCTTGTGTTG  
GTGACCAGCTTTGTTATCGCGGCCTCATCAAGTGTCTAATGAATAAGTAAAACGATT  
25 GCAGT (SEQ ID NO:341)

**Translation:**

MMSILGVLLIICLLCPLTAVLEDGDQPADRPAERMQDGISSEHHFPFLDPVKRCCHLLAC  
RFGCSPCCW (SEQ ID NO:342)

30

**Toxin Sequence:**

Cys-Cys-His-Leu-Leu-Ala-Cys-Arg-Phe-Gly-Cys-Ser-Xaa3-Cys-Cys-Xaa4-^ (SEQ ID NO:343)

35 **Name:** Om3.4  
**Species:** omaria  
**Cloned:** Yes

**DNA Sequence:**

40 CAAGAAGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGTTGATCATCTG  
TCTACTTCTTTGTCCCCTTACTGCTGTTCCGCAGGATGGAGATCAACCTGCAGACCG  
ACCTGCAGAGCGTATGCAGGGCGGCATTTCATCTGAACATCATCCCTTTTTGATCC  
CGTCAAACGGTGTGTCAGGTACGGGTGGACATGCTGGCTAGGATGCACTCCCTGTG  
GTTGTTGACCAGCTTTGTTATCGCGGCCTCATCAAGTGTCTAATGAATAAGTAAAAC  
45 GATTGCAGT (SEQ ID NO:344)

**Translation:**

MMSKLGVLLIICLLLCPLTAVPQDGDQPADRPAERMQGGISSEHHPPFFDPVKRCCRYGW  
TCWLGCTPCGC (SEQ ID NO:345)

**Toxin Sequence:**

5 Cys-Cys-Arg-Xaa5-Gly-Xaa4-Thr-Cys-Xaa4-Leu-Gly-Cys-Thr-Xaa3-Cys-Gly-Cys-^ (SEQ ID NO:346)

10 **Name:** Ep3.1  
**Species:** episcopatus  
**Cloned:** Yes

**DNA Sequence:**

15 CAAGAAGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGTTGACCATCTG  
TCTACTTCTGTTTTCCCTTATTGCTGTTCCGCTTGATGGAGATCAACATGCAGACCAA  
CCTGCAGAGCGTCTGCAGGGCGACATTTTATCTGAAAAGCATCCCTTATTTATGCCT  
GTCAAACGGTGTTGCGATGAGGACGAATGCAACAGTTCATGCTGGCCTTGTTGTTGG  
GGGTGATCAGCTTTGTTATCGCGGCCTGATCAAGTGTATAATGAATAAGTAAAACG  
ATTGCAGT (SEQ ID NO:347)

20

**Translation:**

MMSKLGVLLTICLLLFSLIAVPLDGDQHADQPAERLQGDILSEKHPLFMPVKRCCDEDE  
CNSSCWPCCWG (SEQ ID NO:348)

25 **Toxin Sequence:**

Cys-Cys-Asp-Xaa1-Asp-Xaa1-Cys-Asn-Ser-Ser-Cys-Xaa4-Xaa3-Cys-Cys-Xaa4-# (SEQ ID NO:349)

30 **Name:** Ep3.2  
**Species:** episcopatus  
**Cloned:** Yes

**DNA Sequence:**

35 CAAGAGGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGTTGACCATCTG  
TCTACTTCTGTTTTCCCTTATTGCTGTTCCGCTTGATGGAGATCAACATGCAGACCAA  
CCTGCAGAGCGTCTGCAGGGCGACATTTTATCTGAAAAGCATCCCTTATTTATGCCT  
GTCAAACGGTGTTGCGATGAGGACGAATGCAGCAGTTCATGCTGGCCTTGTTGTTGG  
GGATGAGCAGCTTTGTTATCGCGGCCTCATCAAGTGTCTAATGAATAAGTAAAACG  
40 ATTGCAGT (SEQ ID NO:350)

45 **Translation:**

MMSKLGVLLTICLLLFSLIAVPLDGDQHADQPAERLQGDILSEKHPLFMPVKRCCDEDE  
CSSSCWPCCWG (SEQ ID NO:351)

**Toxin Sequence:**

Cys-Cys-Asp-Xaa1-Asp-Xaa1-Cys-Ser-Ser-Ser-Cys-Xaa4-Xaa3-Cys-Cys-Xaa4-# (SEQ ID NO:352)

5   **Name:**       Ep3.3  
     **Species:**   episcopatus  
     **Cloned:**    Yes

**DNA Sequence:**

10   CAAGAGGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGTTGACCATCTG  
     TCTACTTCTGTTTTCCCTTACTGCTGTTCCGCTTGATGGAGATCAACATGCAGACCAA  
     CCTGCAGAGCGTCTGCAGGGCGACATTTTATCTGAAAAGCATCCCTTATTTAATCCC  
     GTCAAACGGTGTTGCCCGGCGGCGGCATGTGCCATGGGATGCAAGCCTTGTTGTGG  
     ATGAGCAGCTTTGTTATCGTGGCCTCATCAAGTGTCTAATGAATAAGTAAAACGATT  
15   GCAGT (SEQ ID NO:353)

**Translation:**

MMSKLGVLLTICLLLFSLTAVPLDGDQHADQPAERLQGDILSEKHPLFNPVKRCCPAAA  
CAMGCKPCCG (SEQ ID NO:354)

20

**Toxin Sequence:**

Cys-Cys-Xaa3-Ala-Ala-Ala-Cys-Ala-Met-Gly-Cys-Lys-Xaa3-Cys-Cys-# (SEQ ID NO:355)

25   **Name:**       Au3.2  
     **Species:**   aulicus  
     **Cloned:**    Yes

**DNA Sequence:**

30   CAAGAGGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGTTGACCATCTG  
     TCTGCTTCTGTTTTCCGTTACTGCTCTTCCGCCGGATGGAGATCAACCTGCAGACCG  
     AGCTGCAGAGCGTAGGCAGGTCGAGCAGCATCCCGTGTTTGATCATGAAAGAGGGT  
     GTTGCTCGCCACCATGCCACAGTATTTGCGCTGCTTTCTGTTGCGGGTGATGATAAC  
     GTGTTGATGACCCACTTTGTCATCACGGCTGCGTCAAGTGTCTAATGAATAAGTAAA  
35   ATGATTGCAGT (SEQ ID NO:356)

**Translation:**

MMSKLGVLLTICLLLFSVTALPPDGDQPADRAAERRQVEQHPVFDHERGCCSPPCHSIC  
AAFCCG (SEQ ID NO:357)

40

**Toxin Sequence:**

Gly-Cys-Cys-Ser-Xaa3-Xaa3-Cys-His-Ser-Ile-Cys-Ala-Ala-Phe-Cys-Cys-# (SEQ ID NO:358)

45   **Name:**       Au3.3  
     **Species:**   aulicus  
     **Cloned:**    Yes

**DNA Sequence:**

CAAGAGGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGTTGACCATCTG  
TCTACTTCTGTTTTCCCTTACTGCTGTTCCGCTTGATGGAGATCAACATGCAGACCAA  
5 CCTGCAGAGCGTCTGCAGGGCGACATTTTATCTGAAAAGCATCCCTTATTTAATCCC  
GTCAAACGGTGTTGCCGACCGGTGGCATGTGCCATGGGATGCAAGCCTTGTTGTGG  
ATGAGCAGCTTTGTTATCGTGGCCTCATCAAGTGTCTAATGAATAAGTAAAATGATT  
GCAGT (SEQ ID NO:359)

**Translation:**

10 MMSKLGVLITICLLLFSLTAVPLDGDQHADQPAERLQGDILSEKHPLFNPVKRCCRPVA  
CAMGCKPCCG (SEQ ID NO:360)

**Toxin Sequence:**

15 Cys-Cys-Arg-Xaa3-Val-Ala-Cys-Ala-Met-Gly-Cys-Lys-Xaa3-Cys-Cys-# (SEQ ID NO:361)

**Name:** Au3.4

**Species:** aulicus

20 **Cloned:** Yes

**DNA Sequence:**

CAAGAGGGATCGATAGCAGTTCaTGATGTCTAAACTGGGAGTCTTGTTGATCATCTG  
TCTACTTCTGTCTCCCTTACTGCTGTTCCGCTGGATGGAGATCAACCTGCAGACCG  
25 ACCTGCAGAGCGTATGCAGGACGACATTTTCATCTGAACATCAACCCATGTTTGATGC  
CATCAGACAGTGTTGCCCGGCGGTGGCATGCGCCATGGGATGCGAGCCTTGTTGTG  
GATGACCAGCTTTGTTATCGCGGCCTCATCAAGTGTCTAATGAATAAGTAAAATGAT  
TGCAGT (SEQ ID NO:362)

**Translation:**

30 MMSKLGVLIIICLLLSPLTAVPLDGDQPADRP AERMQDDISSEHQPMFDAIRQCCPAVA  
CAMGCEPCCG (SEQ ID NO:363)

**Toxin Sequence:**

35 Xaa2-Cys-Cys-Xaa3-Ala-Val-Ala-Cys-Ala-Met-Gly-Cys-Xaa1-Xaa3-Cys-Cys-# (SEQ ID  
NO:364)

**Name:** Ae3.1

40 **Species:** aureus

**Cloned:** Yes

**DNA Sequence:**

CAAGAAGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGCCTTGTTGACCATCT  
45 GTCTACTTCTGTTTTCCCTTACTGCTGTTCCGCTGGATGGAGATCAACATGCAGACC  
AACATGCAGAGCGTCTGCATGACCGCCTTCCAAGTGAATAATCATCCCTTATATGATC  
CCGTCAAACGGTGTTGCGATGATTCCGAATGCGACTATTCTTGCTGGCCTTGCTGTA

TTTTTGGATAACCTTTGTTATCGCGGCCTCATCAAGTGTCAAATGAATAAGTAAAAC  
GATTGCAGT (SEQ ID NO:365)

**Translation:**

5 MMSKLGALLTICLLLFSLTAVPLDGDQHADQHAERLHDRLPTENHPLYDPVKRCCDDS  
ECDYSCWPCCIFG (SEQ ID NO:366)

**Toxin Sequence:**

10 Cys-Cys-Asp-Asp-Ser-Xaa1-Cys-Asp-Xaa5-Ser-Cys-Xaa4-Xaa3-Cys-Cys-Ile-Phe-# (SEQ ID  
NO:367)

**Name:** Ae3.2

**Species:** aureus

15 **Cloned:** Yes

**DNA Sequence:**

CAAGAGGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGCCTTGTTGACCATCT  
GTCTACTTCTGTTTTCCCTAACTGCTGTTCCGCTGGATGGAGATCAACATGCAGACC  
20 AACCTGCAGAGCGTCTGCAGGACCGCATTCCAACCTGAAAATCATCCCTTATTTGATC  
CGAACAAACGGTGTTGCAATGATTGGGAATGCGACGATTCATGCTGGCCTTGCTGTT  
ATGGATAACCTTTGTTATCGCGGCCTCATCAAGTGTCAAATGAATAAGTAAAACGAT  
TGCAGT (SEQ ID NO:368)

**Translation:**

25 MMSKLGALLTICLLLFSLTAVPLDGDQHADQPAERLQDRIPTENHPLFDPNKRCCNDWE  
CDDSCWPCCYG (SEQ ID NO:369)

**Toxin Sequence:**

30 Cys-Cys-Asn-Asp-Xaa4-Xaa1-Cys-Asp-Asp-Ser-Cys-Xaa4-Xaa3-Cys-Cys-Xaa5-# (SEQ ID  
NO:370)

**Name:** Cn3.1

35 **Species:** consors

**Cloned:** Yes

**DNA Sequence:**

CAAGAGGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGTTGACCATCTG  
40 TTTGCTTCTGTTTCCCTTACTGCTCTTCCAATGGATGGAGATCAATCTGTAGACCGA  
CCTGCAGAGCGTATGCAGGACGACATTTTCATCTGAGCTGCATCCCTTGTTCAATCAG  
AAAAGAATGTGTTGCGGCGAAGGTGCGCCATGCCCCAGCTATTTTCAGAAACAGTCA  
GATTTGTCATTGTTGTTAAATGACAACGTGTCGATGACCAACTTCGTTATCACGACT  
AATGAATAAGTAAAATGATTGCAGT (SEQ ID NO:371)

45

**Translation:**



MMSKLGVLLTICLLLFPLTALPMDGDQSVDRPAERMQDDISSELHPLFNQKRMCCGEG  
APCPSYFRNSQICHCC (SEQ ID NO:372)

**Toxin Sequence:**

- 5 Met-Cys-Cys-Gly-Xaa1-Gly-Ala-Xaa3-Cys-Xaa3-Ser-Xaa5-Phe-Arg-Asn-Ser-Gln-Ile-Cys-His-  
Cys-Cys-^ (SEQ ID NO:373)

10 **Name:** Cn3.3  
**Species:** consors  
**Cloned:** Yes

**DNA Sequence:**

- 15 TAAGAGGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGTTGACCATCTG  
TCTGCTTCTGTTTCCCCTTATTGCTCTTCCAATGGATGGAGATCAACCTGCAGACCGA  
CCTGCAGAGCGTATGCAGGACGACATTTTCATCTCAGCAGCATCCCTTGTTTGATAAG  
AGAGGCCGCTGTTGCGATGTGCCGAACGCATGCTCCGGCAGATGGTGCAGAGATCA  
CGCACAATGTTGCGGATGACGATAACGTGTTGATGACCAACTTTGTGATCACGGCTA  
20 CATCAAGTGAATAAGTAAAACGATTGCAGT (SEQ ID NO:374)

**Translation:**

MMSKLGVLLTICLLLFPLIALPMDGDQPADRP AERMQDDISSQQHPLFDKRGRCCDVPN  
ACSGRWCRDHAQCCG (SEQ ID NO:375)

25 **Toxin Sequence:**

Gly-Arg-Cys-Cys-Asp-Val-Xaa3-Asn-Ala-Cys-Ser-Gly-Arg-Xaa4-Cys-Arg-Asp-His-Ala-Gln-  
Cys-Cys-# (SEQ ID NO:376)

30 **Name:** Cn3.4  
**Species:** consors  
**Cloned:** Yes

**DNA Sequence:**

- 35 CAAGAGGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGTTGACTGTCTG  
TTTGCTTCTGTTTCCCCTTACTGCTCTTCCGATGGATGGAGATCAACCTGCAGACCAA  
CCTGCAGAGCGTATGCAGGACGACATTTTCATCTGAGCAGCATCCCTTGTTTGATAAG  
AGACAAAGGTGTTGCACTGGGAAGAAGGGGTCATGCTCCGGTAAAGCATGCAAAA  
GTCTCAAATGTTGCTCTGGACGATAACGTGTTGATGACCAACTTTGTTATCACGGCT  
40 ACGTCAAGTGTCTAGTGAATAAGTAAAACGATTGCAGT (SEQ ID NO:377)

**Translation:**

- 45 MMSKLGVLLTVCLLLFPLTALPMDGDQPADQPAERMQDDISSEQHPLFDKRQRCCTGK  
KGSCSGKACKSLKCCSGR (SEQ ID NO:378)

**Toxin Sequence:**

Xaa2-Arg-Cys-Cys-Thr-Gly-Lys-Lys-Gly-Ser-Cys-Ser-Gly-Lys-Ala-Cys-Lys-Ser-Leu-Lys-Cys-Cys-Ser-# (SEQ ID NO:379)

5   **Name:**       Em3.1  
      **Species:**   emaciatus  
      **Cloned:**    Yes

**DNA Sequence:**

10   CAAGAGGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGCTGACCATCTGTCTGCTTCTGTT  
      TCCCTTACTGTTCTTCCGATGGATGGAGATCAACCTGCAGACCTACCTGCATTGCGTGCGCAGTTCTT  
      TGACCTGAACATAGTCCCCGTTTGACCCCGTCAAACGGTGCTGCTCGCGGGATTGCAGTGTTCAT  
      CCCTTGTTGCCCGTATGGATCACCTTGATTATTGCGGCCACGTCAAGTGTCTAATGAATAAGTAAATG  
      ATTGCAGT (SEQ ID NO:380)

**Translation:**

MMSKLGVLITICLLLFPLTVLPMGDQPADLPALRAQFFAPEHSPRFDPVKRCCSRDCSVCIPCCPYGSP  
 (SEQ ID NO:381)

20   **Toxin Sequence:**

Cys-Cys-Ser-Arg-Asp-Cys-Ser-Val-Cys-Ile-Xaa3-Cys-Cys-Xaa3-Xaa5-Gly-Ser-Xaa3-^ (SEQ  
 ID NO:382)

25

Where:

Xaa1 is Glu or γ-carboxy-Glu

Xaa2 is Gln or pyro-Glu

Xaa3 is Pro or hydroxy-Pro

30   Xaa4 is Trp or bromo-Trp

Xaa5 is Tyr, <sup>125</sup>I-Tyr, mono-iodo-Tyr, di-iodo-Tyr, O-sulpho-Tyr or O-phospho-Tyr

^ is free carboxyl or amidated C-terminus, preferably free carboxyl

# is free carboxyl or amidated C-terminus, preferably amidated

? = Status of C-term not known.

35

TABLE2

Alignment of μ-Conopeptides (SEQ ID NO:)

TYPE 1

40	A3.4 (F283)	---CCKVQ-CES--C---TPCC^ (383)
	Ak3.1 (F585)	---CCELP-CGPGFC---VPCC^ (384)
	Ar3.1	---CCERP-CNIG-C---VPCC^ (385)
	Bn3.1 (F586)	---CCNWP-CSMG-C---IPCCYY^ (386)
	Bt3.1	---CCELP-CH-G-C---VPCCWP^ (387)
45	Bt3.2	---CCGLP-CN-G-C---VPCCWPS^ (388)
	Bt3.3	---CCSRN-CAV--C---IPCCPNWPA^ (389)
	bt3a	---CCKQS-CTT--C---MPCCW^ (390)

	bt3b	--ACCXQS-CTT--C---MPCC^ (391)
	bt3c	---CCEQS-CTT--C---MPCCW? (392)
	Ca3.3	R--CCRYP-CPDS-C--HGSCCYK^ (393)
	Ca3.4	---CCPPVACNMG-C---KPCC# (394)
5	Ca3.5	---CCDDSECDYS-C---WPCCMF# (395)
	Ca3.6 (F349)	---CCRR--CYMG-C---IPCCF^ (396)
	Circling	---CCPPVACNMG-C---KPCCG^ (397)
	Comatose/Death	SKQCCHLAACRFG-C---TOCCN^ (398)
	Cp3.1 (F594)	S--CCR--DCGED-C---VGCCR^ (399)
10	Ct3.1 (Z726)	---CCDWP-CIPG-C---TPCCLP^ (400)
	Da3.1	---CCDDSECDYS-C---WPCCILS^ (401)
	Da3.2	Z-QCCPPVACNMG-C---EPCC# (402)
	Da3.3	---CCNAGFCRFG-C---TPCCW^ (403)
	Di3.1	Z--CCVHP-C-P--C---TPCCR^ (404)
15	Fi3.1	---CCPWP-CNIG-C---VPCC^ (405)
	Fi3.2	---CCSKN-CAV--C---IPCCP^ (406)
	Fi3.3	---CCRWP-CP-ARC---GSCCL^ (407)
	Fi3.4	---CCELSRCL-G-C---VPCCIS^ (408)
	Fi3.5	---CCELSKCH-G-C---VPCCIP^ (409)
20	Ge3.1 (F590)	Z--CCTF--CNFG-C---QPCCVP^ (410)
	Ge3.2 (F343/Z734)	Z--CCTF--CNFG-C---QPCCLT^ (411)
	Ge3.3 (F590)	Z--CCTF--CNFG-C---QPCCVP^ (412)
	Gm3.1	---CCDDSECDYS-C---WPCCMF# (413)
	Gm3.2	G--CCHLLACRFG-C---SPCCW^ (414)
25	Gm3.3	---CCSWDVCDHPSC---T-CCG# (415)
	La3.1	---CCDWP-CS-G-C---IPCC^ (416)
	Lp3.1 (F340)	ZINCCPWP-CPST-C--RHQCCH^ (417)
	Lv3.1 (F341)	ZINCCPWP-CPDS-C--HYQCCH^ (418)
	Mr3.2	---CCRLS-CGLG-C---HPCC# (419)
30	Mr3.3	--ECCGSFACRFG-C---VPCCV^ (420)
	Mr3.4	SKQCCHLPACRFG-C---TPCCW^ (421)
	Mr3.5 (F286)	-MGCCPFP-CKTS-C--TTLCC# (422)
	Ms3.1 (Z738)	--ACCEQS-CTT--C---FPCC^ (423)
	Nb3.1 (F87)	---CCELP-CGPGFC---VPCC^ (424)
35	Pu3.1 (F339)	---CCN-S-CYMG-C---IPCCF^ (425)
	Qc3.1 (F342)	ZR-CCQWP-CPGS-C---RCCRT# (426)
	Qc3.2	ZR-CCRWP-CPGS-C---RCCRYR^ (427)
	Qc3.3	R--CCRYP-CPDS-C--HGSCCYK^ (428)
	QcIIIA	---CCSQD-CLV--C---IOCCPN# (429)
40	QcIIIB	---CCSRH-CWV--C---IOCCPN? (430)
	Ra3.1 (F351)	Z-TCCS-N-CGED-C---DGCCQ^ (431)
	Scratcher I	---CCR-T-C-FG-C---TOCC# (433)
	Ts3.1 (F592)	---CCH-K-CYMG-C---IPCCI^ (434)
	Ts3.2 (F345)	K--CCRPP-CAMS-C-GMARCCY^ (435)
45	Bt3.5 (Z495)	R--CCRWP-CPSI-C-GMARCCFVMITC^ (436)
	Bt3.6 (Z497)	R--CCRWP-CP-SRC-GMARCCFVMITC^ (437)
	Tx3.1	F--CCDSNWHISDC----ECCY# (438)

	U014	---CCHWNWCDHL-C---SCCGS^ (439)
	U017	--DCCOLPACPFG-C---NOCC# (440)
	U019	---CCAPSACRLG-C---ROCCR^ (441)
	U020	---CCAOSACRLG-C---ROCCR^ (442)
5	U022	---CCAPSACRLG-C---RPCCR^ (443)
	U024	--GCCGSFACRFG-C---VOCCV^ (444)
	U031	---CCSWDVC DHPSC---TCC# (445)
	U032 (F353)	R--CCKFP-CPDS-C--RYLCC# (446)
	Ae3.1	---CCDDSECDYS-C---WPCCIF# (447)
10	Ae3.2	---CCNDWECDDS-C---WPCCY# (448)
	Af3.1	R--CCR-FPCPDT-C---RHLCC# (449)
	Af3.2	---CC--MTC-FG-C---TPCC# (450)
	Af3.3	---CCDDSECDYS-C---WPCCIFS^ (451)
	Af3.4	---CCR-LLC-LS-C---NPCC# (452)
15	Af3.6	---CCDDSECGYS-C---WPCCY# (453)
	Au3.2	G--CCS-PPCHSI-C--AAFCC# (454)
	Au3.3	---CCRPVACAMG-C---KPCC# (455)
	Au3.4	Z--CCPAVACAMG-C---EPCC# (456)
	Em3.1	---CCS-RDC-SV-C---IPCCPYGSP^ (457)
20	Ep3.1	---CCDEDECNSS-C---WPCCW# (458)
	Ep3.2	---CCDEDECSSS-C---WPCCW# (459)
	Ep3.3	---CCPAAACAMG-C---KPCC# (460)
	Om3.1	---CCDEEECSSA-C---WPCCW# (461)
	Om3.3	---CCHLLACRFG-C---SPCCW^ (462)
25	Sf3.1	---CC--PRC-SE-C---NPCC# (463)

TYPE 2

30	Pn3.2 (AA049)	-RCC--KFP-CPDS-C--KYLCC# (464)
	Fd3.2 (Z831)	-RCC--RWP-CPSI-C-GMARCCSS^ (465)
	Pu3.3 (AA405)	--CC--KLL-CYSG-C---TPCCHI^ (466)
	Eb3.1 (Z821)	--CC--EQP-CYMG-C---IPCCF^ (467)
	Eb3.2 (Z822)	--CC--AQP-CYMG-C---IPCCF^ (468)
35	Pu3.2 (AA403)	--CC--V-S-CYMG-C---IPCCF^ (469)
	Mf3.1 (Z882)	--CC--DWP-CSAG-C---YPCCFP^ (470)
	Mf3.2 (Z885)	-GCC--PPM-C-TP-C---FPCCFR^ (471)
	Ra3.2 (AA414)	RGCCAPPRK-CKDRACK-PARCCGP# (472)
	Sm3.3 (AA419)	ZRCCNGRRG-CSSRWCRDHSRCC# (473)
40	Cn3.3	GRCCDVPNA-CSGRWCRDHAQCC# (474)
	Cn3.4	ZRCCTGKKGSCSGKACKSL-KCCS# (475)

TYPE 3

45	A3.1	-MCCGEGRKCPSYFRNSQICHCC^ (476)
	A3.2 (F84)	--CCR--WPCPRQIDGEY-CGCCL# (477)
	Bu3.5	-RCCGEGLTCPRYWKNSQICACC^ (478)
	Ca3.1	--CCGPGGSCP VYFRDNFICGCC^ (479)

	Cr3.1	RKCCGKDGPCPKYFKDNFICGCC^ (480)
	E3.1	--CCS--WPCPRYSNGKLVCFCL# (481)
	M3.2	--CCGPGGSCPVIYFRDNFICGCC^ (482)
	M3.3	-MCCGESAPCPSYFRNSQICHCC^ (483)
5	M3.4	ZKCCGPGGSCPVIYFTDNFICGCC^ (484)
	M3.5	ZKCCGPGGSCPVIYFRDNFICGCC^ (485)
	S3.1	ZKCCGEGSSCPKYFKNNFICGCC^ (486)
	U001	ZKCCS--GGSCPLYFRDRLICPCC^ (487)
	U034	ZKCCGPGASCPRYFKDNFICGCC^ (488)
10	Cn3.1	-MCCGEGAPCPSYFRNSQICHCC^ (489)

TYPE 4

15	A3.3 (F83)	ZK--CCTGK---KGSCSGKACKNL-KCCS# (490)
	A3.5 (Z488)	ZK--CCTGR---KGSCSGKACKNL-KCCS# (491)
	Bu3.1	VTDRCK----GKREC--GRWCRDHSRCC# (492)
	Bu3.1A	VGDRCK----GKRGK--GRWCRDHSRCC# (493)
	Bu3.2	VGERCK---NGKRGK--GRWCRDHSRCC# (494)
20	Bu3.3	IVDRCCN-KGNGKRGK--SRWCRDHSRCC# (495)
	Bu3.4	VGLYCCRPKPNGQMMC--DRWCEKNSRCC# (496)
	Ca3.2	-RD-CCTPP---KK-CKDRQCKPQ-RCCA# (497)
	L3.1	GRD-CCTPP---RK-CKDRACKPQ-RCCG# (498)
	L3.2	ZRL-CCGFP---KS-CRSRQCKPH-RCC# (499)
25	La3.2	-RD-CCTPP---KK-CRDRQCKPA-RCCG# (500)
	La3.3	RPP-CCTYD---GS-CLKESCMRK-ACC# (501)
	La3.3A	RPP-CCTYD---GS-CLKESCKRK-ACC# (502)
	$\mu$ -GIIIA	-RD-CCTOO---KK-CKDRQCKOQ-RCCA# (503)
	$\mu$ -GIIIB	-RD-CCTOO---RK-CKDRRCKOM-KCCA# (504)
30	$\mu$ -GIIIC	-RD-CCTOO---KK-CKDRRCKOL-KCCA# (505)
	$\mu$ -PIIIA	ZRL-CCGFO---KS-CRSRQCKOH-RCC# (506)
	M3.1	-RD-CCTPP---KK-CKDRQCKPQ-RCCA# (507)
	Mr3.1	RGG-CCTPP---RK-CKDRACKPA-RCCGP# (508)
	Nb3.2 (F582)	ZK--CCTGK---KGSCSGKACKNL-KCCS# (509)
35	Pr3.1 (Z500)	RGG-CCTPP---KK-CKDRACKPA-RCCGP# (510)
	Pr3.2 (Z501)	-RG-CCTPP---RK-CKDRACKPA-RCCGP# (511)
	R3.1	LOS-CCSLN---LRLCOVOACKRN-OCCT# (512)
	R3.2	ZQR-CCTVK----RICOVOACRSK-OCCKS^ (513)
	R3.3	RGG-CCTPP---RK-CKDRACKPA-RCCGP# (514)
40	Sm3.1	ZK--CCTGK---KGSCSGKACKNL-KCCS# (515)
	T3.1	H-G-CCKGO---EG-CSSRECROQ-HCC# (516)
	T3.2 (Y088)	H-G-CCEGP---KG-CSSRECRPQ-HCC# (517)
	Wi3.1 (M548)	LPS-CCDFE----RLCVVPACIRH-QCCT# (518)

45

Type 5

Om3.2	CCKYGWTCLLGCTPCDC^ (519)
-------	--------------------------

Om3.4

CCRYGWTGWLGCTPCGC<sup>^</sup> (520)Type 6

5 S3.2 (F352)

Z-NCCNGG-CSSKWCRDHARCC# (432)

## EXAMPLE 3

Effect of Intrathecal Administration of  $\mu$ -Conopeptides

10 [0087] Male C57 black mice (20-25g) are obtained from Charles River Laboratories. These mice and the animals are housed in a temperature controlled ( $23^{\circ} \pm 3^{\circ} \text{C}$ ) room with a 12 hour light-dark cycle with free access to food and water. All animals are euthanized in accordance with Public Health Service policies on the humane care of laboratory animals.

15 [0088] Intrathecal (it) drug injections are performed as described (Hylden and Wilcox, 1980). A  $\mu$ -conopeptide or vehicle is administered in a volume of 5  $\mu\text{l}$ . Duration of hind-limb paralysis is assessed. This experiment reveals that injection of  $\mu$ -conopeptides into the intrathecal space of C57 black mice produced a paralysis of the animal. The animals in this experiment recovered fully.

20

## EXAMPLE 4

Effect of  $\mu$ -Conopeptides as a Local Anesthetic

25 [0089] Male Hartley guinea pigs (retired breeders) are obtained from Charles River Laboratories. The local anesthetic test is performed essentially as described (Bulbring and Wajda, 1945). On the day prior to test day, a patch on the back of the guinea pig is denuded of hair, first by shaving with electric clippers and subsequently with depilatory cream (Nair®). Depilatory cream is applied for five minutes and removed with a warm washcloth. The guinea pigs are dried and returned to their cages. On the following day, intradermal injections (0.1 ml vols) of lidocaine, bupivacaine, a  $\mu$ -conopeptide or vehicle (0.5% cyclodextran) are made into the denuded patch. The injection produced a raised wheal on the surface of the skin which is 30 circled with a felt-tipped pen. Typically, four injections are made on the back of each guinea pig. In some cases, guinea pigs are reused following at least one week of recovery and injecting into an unused portion of the skin. The stimulus consists of mild pin pricks (not hard enough to break the skin) with a 26G needle. The response is a localized skin twitch caused by contraction

of cutaneous muscles. A unit test consisted of six uniform pin pricks, 3-5 seconds apart, within the injected area. Unit scores range from 0 (complete anesthesia) to 6 (no anesthesia). For potency experiments, the unit test is repeated at each site at five minute intervals for 30 minutes, and unit test scores summed (with 36 representing no anesthesia to 0 representing complete anesthesia. For duration experiments, unit tests are performed as described over the course of several hours to days.

[0090]  $\mu$ -Conopeptides of the present invention produce a potent and long lasting local anesthetic effect in the intracutaneous wheal test in the guinea pig. As expected, bupivacaine has a slightly longer duration than lidocaine, consistent with clinical observations.

10

#### EXAMPLE 5

##### Muscle Relaxant Effect of $\mu$ -Conopeptides in Anesthetized Monkeys

[0091]  $\mu$ -Conopeptides are dissolved 0.9 percent saline at a concentration of 2 mg/ml. Rhesus monkeys are anesthetized with halothane, nitrous oxide and oxygen. The maintenance concentration of halothane is 1.0%. Arterial and venous catheters are placed in the femoral vessels for drug administration and recording of the arterial pressure. Controlled ventilation is accomplished via an endotracheal tube. Twitch and tetanic contractions of the tibialis anterior muscle are elicited indirectly via the sciatic nerve. Recordings of arterial pressure, electrocardiogram (lead I), heart rate, and muscle function are made simultaneously. Four to six animals received each listed compound. Four additional animals received succinylcholine chloride or d-tubocurarine chloride as controls. It is seen that the tested  $\mu$ -conopeptides generally provide similar or better results than those seen for succinylcholine chloride or d-tubocurarine chloride.

25

#### EXAMPLE 6

##### *In vivo* Activity of $\mu$ -Conopeptides in Pain Models

[0092] The anti-pain activity of  $\mu$ -conopeptides is shown in several animal models. These models include the nerve injury model (Chaplan, et al., 1997), the nociceptive response to s.c. formalin injection in rats (Codene, 1993) and an NMDA-induced persistent pain model (Liu, et al., 1997). In each of these models it is seen that the  $\mu$ -conopeptides and  $\mu$ -conopeptides derivatives have analgesic properties.

30

[0093] More specifically, this study evaluates the effect of intrathecal administration of  $\mu$ -conopeptides in mice models of nociceptive and neuropathic pain. For nociceptive pain, the effect of the  $\mu$ -conopeptides is studied in two different tests of inflammatory pain. The first is the formalin test, ideal because it produces a relatively short-lived, but reliable pain behavior that is readily quantified. There are two phases of pain behavior, the second of which is presumed to result largely from formalin-evoked inflammation of the hind paw. A  $\mu$ -conopeptide is administered 10 minutes prior to injection of formalin. The number of flinches and/or the duration of licking produced by the injection is monitored. Since the first phase is presumed to be due to direct activation of primary afferents, and thus less dependent on long term changes in the spinal cord,  $\mu$ -conopeptides are presumed to have greatest effect on the magnitude of pain behavior in the second phase.

[0094] The mechanical and thermal thresholds in animals that received an injection of complete Freund's adjuvant into the hind paw are also studied. This produces a localized inflammation including swelling of the hind paw and a profound decrease in mechanical and thermal thresholds, that are detected within 24 hours after injection. The changes in thresholds in rats that receive  $\mu$ -conopeptides are compared with those of rats that receive vehicle intrathecal injections.

[0095] An important issue is whether the drugs are effective when administered after the pain model has been established, or whether they are effective only if used as a pretreatment. Clearly, the clinical need is for drugs that are effective after the pain has developed. To address this issue, animals are studied in which  $\mu$ -conopeptides are administered repeatedly, after the inflammation (CFA) or nerve injury has been established. In these experiments, a  $\mu$ -conopeptide is injected daily by the intrathecal (i.t.) route. The mechanical and thermal thresholds (measured, respectively, with von Frey hairs in freely moving animals and with the Hargreave's test, also in freely moving animals) are repeated for a 2 to 4 week period after the injury is induced and the changes in pain measured monitored over time.

#### EXAMPLE 7

##### Effect of $\mu$ -Conotoxins in a Pain Model

[0096] Analgesic activity of  $\mu$ -conotoxins is also tested in pain models as follows.

[0097] Persistent pain (formalin test). Intrathecal (it) drug injections are performed as described by Hylden and Wilcox (1980). An  $\mu$ -conopeptide or vehicle is administered in a



volume of 5  $\mu$ l. Fifteen minutes after the i.t. injection, the right hindpaw is injected with 20  $\mu$ l of 5% formalin. Animals are placed in clear plexiglass cylinders backed by mirrors to facilitate observation. Animals are closely observed for 2 minutes per 5 minute period, and the amount of time the animal spent licking the injected paw is recorded in this manner for a total of 45-50  
5 minutes. Results are expressed as licking time in seconds per five minutes. At the end of the experiment, all animals are placed on an accelerating rotorod and the latency to first fall was recorded.  $\mu$ -Conopeptides are found to be active in this model which is predictive of efficacy for treating neuropathic pain.

[0098] Acute pain (tail-flick). A  $\mu$ -conopeptide or saline is administered intrathecally  
10 (i.t.) according to the method of Hylden and Wilcox (1980) in a constant volume of 5  $\mu$ l. Mice are gently wrapped in a towel with the tail exposed. At various time-points following the i.t. injection, the tail is dipped in a water bath maintained at 54° C. and the time to a vigorous tail withdrawal is recorded. If there is no withdrawal by 8 seconds, the tail is removed to avoid tissue damage.

15 [0099] Neuropathic pain. The partial sciatic nerve ligation model is used to assess the efficacy of  $\mu$ -conopeptides in neuropathic pain. Nerve injury is produced according to the methods of Malmberg and Basbaum (1998). Animals are anesthetized with a ketamine/xylazine solution, the sciatic nerve is exposed and tightly ligated with 8-0 silk suture around 1/3 to 1/2 of the nerve. In sham-operated mice the nerve is exposed, but not ligated. Animals are allowed to  
20 recover for at least 1 week before testing is performed. On the testing day, mice are placed in plexiglass cylinders on a wire mesh frame and allowed to habituate for at least 60 minutes. Mechanical allodynia is assessed with calibrated von Frey filaments using the up-down method as described by Chaplan et al. (1994), and the 50% withdrawal threshold is calculated. Animals that did not respond to any of the filaments in the series are assigned a maximal value of 3.6  
25 grams, which is the filament that typically lifted the hindlimb without bending, and corresponds to approximately 1/10 the animal's body weight.

[0100] The data obtained demonstrate that  $\mu$ -conopeptides have potent analgesic properties in three commonly used models of pain: acute, persistent/inflammatory and neuropathic pain models.

## EXAMPLE 8

Activity of  $\mu$ -Conopeptide S3.2 on Neuronal Sodium Channels

[0101]  $\mu$ -Conopeptide S3.2 was tested for activity on sodium channels as follows. S3.2 was administered to mice by intracerebroventricular (ICV) injection. Administration of S3.2 in this manner caused mice to show a spectrum of activity that is characteristic of all sodium channel blockers, including rapid loss of righting reflex, coma-like inactivity and spastic uncontrolled limb movement. Following intrathecal (it) administration to mice, S3.2 causes rapid hindlimb paralysis that spreads to include the entire body over a course of 10-20 minutes followed by death, presumably due to respiratory paralysis. However, unlike classic  $\mu$ -conopeptides, S3.2 has no significant activity following intravenous administration (iv) to mice. Classic  $\mu$ -conopeptides, such as GIIIA and PIIIA, cause rapid paralysis and death following iv administration, indicating their activity at skeletal muscle sodium channels. To confirm the selectivity of S3.2, 80 nmol was administered iv to rats. The effect of S3.2 was measured on skeletal muscle contraction, blood pressure and heart rate. S3.2 was found to have no effect on any of these parameters. Controls were performed using classical  $\mu$ -conopeptides, including Sm3.1, Sm3.3 and Bu3.1 described herein, also administered iv at 80 nmol. These control peptides caused a dramatic decrease in skeletal muscle contractility, as well as a significant drop in systemic blood pressure. Thus,  $\mu$ -conopeptide S3.2 surprisingly is selective for neuronal sodium channels. The most obvious difference between the S3.2 sequence and the sequences of these other peptides is a shortened first loop (the first loop between cysteine residues) which lacks a charged amino acid.

[0102] It will be appreciated that the methods and compositions of the instant invention can be incorporated in the form of a variety of embodiments, only a few of which are disclosed herein. It will be apparent to the artisan that other embodiments exist and do not depart from the spirit of the invention. Thus, the described embodiments are illustrative and should not be construed as restrictive.

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WHAT IS CLAIMED IS:

1. An isolated peptide selected from the group consisting of:
  - (a) a peptide set forth in Table 1 or Table 2 and
  - (b) a derivative of the peptide in (a).
2. The isolated peptide of claim 1, wherein Xaa<sub>1</sub> is Glu, Xaa<sub>2</sub> is pyro-Glu, Xaa<sub>4</sub> is Trp and Xaa<sub>5</sub> is Tyr.
3. The derivative of the peptide of claim 1, in which the Arg residues may be substituted by Lys, ornithine, homoargine, nor-Lys, N-methyl-Lys, N,N-dimethyl-Lys, N,N,N-trimethyl-Lys or any synthetic basic amino acid; the Lys residues may be substituted by Arg, ornithine, homoargine, nor-Lys, or any synthetic basic amino acid; the Tyr residues may be substituted with meta-Tyr, ortho-Tyr, nor-Tyr, mono-halo-Tyr, di-halo-Tyr, O-sulpho-Tyr, O-phospho-Tyr, nitro-Tyr or any synthetic hydroxy containing amino acid; the Ser residues may be substituted with Thr or any synthetic hydroxylated amino acid; the Thr residues may be substituted with Ser or any synthetic hydroxylated amino acid; the Phe residues may be substituted with any synthetic aromatic amino acid; the Trp residues may be substituted with Trp (D), neo-Trp, halo-Trp (D or L) or any aromatic synthetic amino acid; the Asn, Ser, Thr or Hyp residues may be glycosylated; the Tyr residues may also be substituted with the 3-hydroxyl or 2-hydroxyl isomers (meta-Tyr or ortho-Tyr, respectively) and corresponding O-sulpho- and O-phospho-derivatives; the acidic amino acid residues may be substituted with any synthetic acidic amino acid, e.g., tetrazolyl derivatives of Gly and Ala; the aliphatic amino acids may be substituted by synthetic derivatives bearing non-natural aliphatic branched or linear side chains C<sub>n</sub>H<sub>2n+2</sub> up to and including n=8; the Met residues may be substituted by Nle; the Cys residues may be in D or L configuration and may optionally be substituted with homocysteine (D or L); pairs of Cys residues may be replaced pairwise with isoteric lactam or ester-thioether replacements, such as Ser/(Glu or Asp), Lys/(Glu or Asp), Cys/Glu (or Asp) or Cys/Ala combinations; and the peptide may be radioiodinated or radiotriated.

4. A substantially pure  $\mu$ -conotoxin peptide derivative comprising a permutant of the peptide of claim 1,2 or 3.
5. A substantially pure  $\mu$ -conotoxin peptide derivative comprising the peptide or peptide derivative of claim 1, 2 or 3 modified to contain an O-glycan, an S-glycan or an N-glycan.
6. A substantially pure  $\mu$ -conotoxin peptide derivative comprising the peptide derivative of claim 4 modified to contain an O-glycan, an S-glycan or an N-glycan.
7. An isolated nucleic acid encoding a  $\mu$ -conopeptide propeptide having an amino acid sequence set forth in Table 1.
8. The isolated nucleic acid of claim 7, wherein the nucleic acid comprises a nucleotide sequence set forth in Table 1.
9. An isolated  $\mu$ -conopeptide propeptide having an amino acid sequence set forth in Table 1.
10. A method for treating or preventing disorders associated with voltage gated ion channel disorders in which comprises administering to a patient in need thereof a therapeutically effective amount of a peptide of claim 1 or a pharmaceutically acceptable salt thereof.
11. The method of claim 10, wherein said disorder is a neurologic disorder.
12. The method of claim 11, wherein said neurologic disorder is Amytrophic Lateral Sclerosis.
13. The method of claim 11, wherein said neurologic disorder is head trauma.
14. The method of claim 11, wherein said neurologic disorder is epilepsy.

15. The method of claim 11, wherein said neurologic disorder is a neurotoxic injury associated with conditions of hypoxia, anoxia or ischemia.
16. The method of claim 15, wherein said neurotoxic injury is associated with stroke,  
5 cerebrovascular accident, brain or spinal cord trauma, myocardial infarct, physical trauma, drownings, suffocation, perinatal asphyxia, or hypoglycemic events.
17. The method of claim 10, wherein said disorder is pain.
- 10 18. The method of claim 17, wherein said pain is migraine, acute pain, persistent pain, chronic pain, neuropathic pain or nociceptive pain.
19. The method of claim 18, wherein the pain is trigeminal neuralgia, diabetic neuropathy, post-herpetic neuralgia, neuroma pain, phantom limb pain.
- 15 20. The method of claim 17, wherein said pain is burn pain.
21. The method of claim 10, wherein said disorder is a neuromuscular disorder.
- 20 22. The method of claim 21, wherein said neuromuscular disorder is myofacial pain syndrome, chronic muscle spasm, dystonias or spasticity.
23. A method for providing musculoskeletal relaxation in a patient undergoing a surgical  
25 procedure requiring anesthesia which comprises administering to a patient in need thereof a therapeutically effective amount of a peptide of claim 1 or a pharmaceutically acceptable salt thereof.
24. A method of alleviating pain which comprises administering to a mammal that is either  
30 exhibiting pain or is about to be subjected to a pain-causing event a pain-alleviating amount of a peptide of claim 1 or a pharmaceutically acceptable salt thereof.
25. The method of claim 24, wherein the peptide is administered as a local anesthetic.



26. The method of claim 24, wherein the peptide is administered as an ocular anesthetic.
27. A method for characterizing a pore occlusion site on a sodium channel subtype  
5 comprising determining the affinity of said site for a peptide of claim 1.
28. The method of claim 27, wherein said sodium channel subtype is a neuronal sodium  
channel subtype and said peptide is  $\mu$ -conopeptide S3.2 comprising an amino acid  
sequence as set forth in SEQ ID NO:211 or SEQ ID NO:432.
- 10 29. A method for screening a small molecule library to identify a small molecule which is a  
selective blocking agent of a sodium channel subtype comprising (a) measuring the  
blocking activity of a small molecule on said sodium channel subtype, (b) measuring the  
blocking activity of a peptide of claim 1 on said sodium channel subtype and (c)  
15 comparing the blocking activity of said small molecule with the blocking activity of said  
peptide.
30. The method of claim 29, wherein said sodium channel subtype is a neuronal sodium  
channel subtype and said peptide is  $\mu$ -conopeptide S3.2 comprising an amino acid  
20 sequence as set forth in SEQ ID NO:211 or SEQ ID NO:432.
31. A method for screening a small molecule library to identify a small molecule which is a  
selective blocking agent of a sodium channel subtype comprising (a) measuring the  
binding affinity of a small molecule on said sodium channel subtype, (b) measuring the  
25 binding affinity of a peptide of claim 1 on said sodium channel subtype and (c)  
comparing the binding affinity of said small molecule with the binding affinity of said  
peptide.
32. The method of claim 31, wherein said peptide is radiolabeled.

33. The method of claim 31, wherein said sodium channel subtype is a neuronal sodium channel subtype and said peptide is  $\mu$ -conopeptide S3.2 comprising an amino acid sequence as set forth in SEQ ID NO:211 or SEQ ID NO:432.
- 5 34. The method of claim 33, wherein said peptide is radiolabeled.
35. A method for screening a small molecule library to identify a small molecule which is a selective blocking agent of a sodium channel subtype comprising (a) allowing a peptide of claim 1 to bind to a sodium channel subtype, (b) adding a small molecule and (c)  
10 measuring the amount of displacement of said peptide on said sodium channel subtype by said small molecule.
36. The method of claim 35, wherein said peptide is radiolabeled.
- 15 37. The method of claim 35, wherein said sodium channel subtype is a neuronal sodium channel subtype and said peptide is  $\mu$ -conopeptide S3.2 comprising an amino acid sequence as set forth in SEQ ID NO:211 or SEQ ID NO:432.
38. The method of claim 37, wherein said peptide is radiolabeled.
- 20 39. A method for screening a small molecule library to identify a small molecule which is a selective blocking agent of a sodium channel subtype comprising (a) allowing a small molecule to bind to a sodium channel subtype, (b) adding a peptide of claim 1 and (c) measuring the amount of displacement of said small molecule on said sodium channel  
25 subtype by said small peptide.
40. The method of claim 39, wherein said sodium channel subtype is a neuronal sodium channel subtype and said peptide is  $\mu$ -conopeptide S3.2 comprising an amino acid sequence as set forth in SEQ ID NO:211 or SEQ ID NO:432.
- 30 41. A method of identifying compounds that mimic the therapeutic activity of a  $\mu$ -conotoxin, comprising the steps of: (a) conducting a biological assay on a test compound to

determine the therapeutic activity; and (b) comparing the results obtained from the biological assay of the test compound to the results obtained from the biological assay of a  $\mu$ -conotoxin, wherein said  $\mu$ -conotoxin is a peptide of claim 1.

- 5    42.    The method of claim 41, wherein said  $\mu$ -conotoxin is S3.2 comprising an amino acid set forth in SEQ ID NO:211 or SEQ IN NO:432.

## SEQUENCE LISTING

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agagcgtatg caggacgact ttataactga gcatcatccc ctgtttgatc ctgtcaaacg 180

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Ala Glu Arg Met Gln Asp Asp Phe Ile Thr Glu His His Pro Leu Phe

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atttcacatctg atcaacatct cttctttgat ctcacaaac ggtgctgcga gttgccatgc 180  
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20 25 30  
Arg Pro Ala Glu Arg Met Gln Asp Ile Ser Ser Asp Gln His Leu Phe  
35 40 45  
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50 55 60  
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3

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 gtgttgcggc gaaggccgga aatgccccag ctatttcaga aacagtcaga tttgtcattg 240  
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 20 25 30  
 Glu Glu Arg Met Gln Asp Asp Ile Ser Ser Glu Gln His Pro Leu Phe  
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Cys Cys Leu

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 20 25 30  
 Gln Pro Ala Asp Arg Met Gln Asp Asp Ile Ser Ser Glu Gln Tyr Pro  
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 Arg His Ala Glu Arg Met His Asp Gly Ile Ser Pro Lys Arg His Pro  
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 s Pro or Hyp

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 actgctcttc cgatggatgg agatcaacct gcagaccgac ctgcagagcg tagtcaggac 120  
 gtttcatctg aacagcatcc cttgtttgat cccgtcaaac ggtgttgcaa ctggccatgc 180  
 tccatgggat gcattcccttg ttgctactat taataacgtg ttgatgacca actttctcga 240  
 g 241  
 <210> 20  
 <211> 70  
 <212> PRT  
 <213> Conus bandus

<400> 20  
 Gly Ser Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Met Leu  
 1 5 10 15  
 Leu Phe Pro Leu Thr Ala Leu Pro Met Asp Gly Asp Gln Pro Ala Asp  
 20 25 30  
 Arg Pro Ala Glu Arg Ser Gln Asp Val Ser Ser Glu Gln His Pro Leu  
 35 40 45  
 Phe Asp Pro Val Lys Arg Cys Cys Asn Trp Pro Cys Ser Met Gly Cys  
 50 55 60

Ile Pro Cys Cys Tyr Tyr  
65 70

<210> 21  
<211> 16  
<212> PRT  
<213> Conus bandus

<220>  
<221> PEPTIDE  
<222> (1)..(16)  
<223> Xaa at residue 5 and 12 is Pro or Hyp; Xaa at residue 4 is Trp or bromo-Trp; Xaa at residue 15 and 16 is Tyr, 125I-Tyr, mono-iodo-Tyr, di-iodo-Tyr, O-sulpho-Tyr or O-phospho-Ty

<400> 21  
Cys Cys Asn Xaa Xaa Cys Ser Met Gly Cys Ile Xaa Cys Cys Xaa Xaa  
1 5 10 15

<210> 22  
<211> 298  
<212> DNA  
<213> Conus betulinus

<400> 22  
caagagggat cgatagcagt tcatgatgtc taaactggga gtcttggtga ccttctgtot 60  
gcttctgttt cccctgactg ctcttccgct ggatgaagat caacctgcag accgacctgc 120  
agagcgtatg caggacattt catctgaaca gcatcccttg ttgatcccg tcaaacggtg 180  
ttgcgaattg ccatgccatg gatgcgtccc ttgttgctgg ccttaataac gtgtggatga 240  
ccaactgtgt tatcacggcc acgtcaagtg tctaataaat aagtaaaatg attgcagt 298

<210> 23  
<211> 67  
<212> PRT  
<213> Conus betulinus

<400> 23  
Met Met Ser Lys Leu Gly Val Leu Leu Thr Phe Cys Leu Leu Leu Phe  
1 5 10 15

Pro Leu Thr Ala Leu Pro Leu Asp Glu Asp Gln Pro Ala Asp Arg Pro  
20 25 30

Ala Glu Arg Met Gln Asp Ile Ser Ser Glu Gln His Pro Leu Phe Asp  
35 40 45

Pro Val Lys Arg Cys Cys Glu Leu Pro Cys His Gly Cys Val Pro Cys  
50 55 60

Cys Trp Pro  
65

<210> 24  
<211> 15  
<212> PRT  
<213> Conus betulinus

<220>  
<221> PEPTIDE  
<222> (1)..(15)  
<223> Xaa at residue 3 is Glu or gamma-carboxy Glu; Xaa at residue 5, 1

1 and 15 is Pro or Hyp; Xaa at residue 14 is Trp or bromo-Trp

<400> 24

Cys Cys Xaa Leu Xaa Cys His Gly Cys Val Xaa Cys Cys Xaa Xaa  
1 5 10 15

<210> 25

<211> 298

<212> DNA

<213> Conus betulinus

<400> 25

caagagggat cgatagcagt tcatgatgtc taaactggga gtcttggtga ccttctgtct 60  
gcttctgttt cccctgactg ctcttcc\_t ggatgaagat caacctgcag accgacatgc 120  
agagcgtatg caggacattt cacctgaaca gcatccctcg ttgatcccg tcaaacgggtg 180  
ttgcgggctg ccatgcaatg gatgcgtccc ttgttgctgg ccttcataac gtgtggacga 240  
ccaactttgt tatcacggcc acgtcaagtg tctgatgaat aagtaaacg attgcagt 298

<210> 26

<211> 68

<212> PRT

<213> Conus betulinus

<400> 26

Met Met Ser Lys Leu Gly Val Leu Leu Thr Phe Cys Leu Leu Leu Phe  
1 5 10 15

Pro Leu Thr Ala Leu Pro Leu Asp Glu Asp Gln Pro Ala Asp Arg His  
20 25 30

Ala Glu Arg Met Gln Asp Ile Ser Pro Glu Gln His Pro Ser Phe Asp  
35 40 45

Pro Val Lys Arg Cys Cys Gly Leu Pro Cys Asn Gly Cys Val Pro Cys  
50 55 60

Cys Trp Pro Ser  
65

<210> 27

<211> 16

<212> PRT

<213> Conus betulinus

<220>

<221> PEPTIDE

<222> (1)..(16)

<223> Xaa at residue 5, 11 and 15 is Pro or Hyp; Xaa at residue 14 is Trp or bromo-Trp

<400> 27

Cys Cys Gly Leu Xaa Cys Asn Gly Cys Val Xaa Cys Cys Xaa Xaa Ser  
1 5 10 15

<210> 28

<211> 282

<212> DNA

<213> Conus betulinus

<400> 28

caagagggat cgatagcagt tcatgatgtt taaactggga gtcttggtga ccatctatat 60

9

gcttctgttt ccctttactg ctcttccgct ggatggagat caacctgcag accaacctct 120  
 agagcgcacg cagtatgaca tggtacgtgc agtgaatccc tggtttgatc ccgtcaaaag 180  
 gtgctgctcg aggaactgcg cagtatgcat cccttggtgc ccgaattggc cagcttgatt 240  
 atcgcgccca agagtctaata gaataagtaa aacgattgca gt 282

<210> 29  
 <211> 71  
 <212> PRT  
 <213> Conus betulinus

<400> 29  
 Met Met Phe Lys Leu Gly Val Leu Leu Thr Ile Tyr Met Leu Leu Phe  
 1 5 10 15  
 Pro Phe Thr Ala Leu Pro Leu Asp Gly Asp Gln Pro Ala Asp Gln Pro  
 20 25 30  
 Leu Glu Arg Met Gln Tyr Asp Met Leu Arg Ala Val Asn Pro Trp Phe  
 35 40 45  
 Asp Pro Val Lys Arg Cys Cys Ser Arg Asn Cys Ala Val Cys Ile Pro  
 50 55 60  
 Cys Cys Pro Asn Trp Pro Ala  
 65 70

<210> 30  
 <211> 18  
 <212> PRT  
 <213> Conus betulinus

<220>  
 <221> PEPTIDE  
 <222> (1)..(18)  
 <223> Xaa at residue 11, 14 and 17 is Pro or Hyp; Xaa at residue 16 is  
 Trp or bromo-Tr

<400> 30  
 Cys Cys Ser Arg Asn Cys Ala Val Cys Ile Xaa Cys Cys Xaa Asn Xaa  
 1 5 10 15

Xaa Ala

<210> 31  
 <211> 325  
 <212> DNA  
 <213> Conus bullatus

<400> 31  
 caagaaggat cgatagcagt tcatgatgtc taaactggga gtcttggtga ccatctgtct 60  
 gcttctgttt cccctttttg ctcttccgca ggatggagat caacctgcag accgacctgc 120  
 agagcgtatg caggacgaca tttcatctga gcagaattcc ttgcttgaga agagagttac 180  
 tgacaggtgc tgcaaaggga agagggaatg cggcagatgg tgcagagatc actcgcgttg 240  
 ttgcggtcga cgataagctg ttgatgacca gctttgttat cacggctaca tcaagtgtct 300  
 agtgaataag taaaatgatt gcagt 325  
 <210> 32

10

<211> 77  
 <212> PRT  
 <213> Conus bullatus

<400> 32  
 Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15  
 Pro Leu Phe Ala Leu Pro Gln Asp Gly Asp Gln Pro Ala Asp Arg Pro  
 20 25 30  
 Ala Glu Arg Met Gln Asp Asp Ile Ser Ser Glu Gln Asn Ser Leu Leu  
 35 40 45  
 Glu Lys Arg Val Thr Asp Arg Cys Cys Lys Gly Lys Arg Glu Cys Gly  
 50 55 60  
 Arg Trp Cys Arg Asp His Ser Arg Cys Cys Gly Arg Arg  
 65 70 75

<210> 33  
 <211> 23  
 <212> PRT  
 <213> Conus bullatus

<220>  
 <221> PEPTIDE  
 <222> (1)..(23)  
 <223> Xaa at residue 11 is Glu or gamma-carboxy Glu; Xaa at residue 15  
 is Trp or bromo-Trp

<400> 33  
 Val Thr Asp Arg Cys Cys Lys Gly Lys Arg Xaa Cys Gly Arg Xaa Cys  
 1 5 10 15  
 Arg Asp His Ser Arg Cys Cys  
 20

<210> 34  
 <211> 326  
 <212> DNA  
 <213> Conus bullatus

<400> 34  
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 agagcgtatg caggatgaca ttcatctga gcagaatccc ttgcttgaga agagagttgg 180  
 tgacaggtgc tgcaaaggga agagggggtg cggcagatgg tgcagagatc actcacgttg 240  
 ttgcggtcga cgataacgtg ttgatgacca gctttgttat cacggctaca tcaagtgtct 300  
 tagtgattaa gtaaaacgat tgcagt 326

<210> 35  
 <211> 77  
 <212> PRT  
 <213> Conus bullatus

<400> 35  
 Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15

11

Pro Leu Phe Ala Leu Arg Gln Asp Gly Asp Gln Pro Ala Asp Arg Pro  
20 25 30

Ala Glu Arg Met Gln Asp Asp Ile Ser Ser Glu Gln Asn Pro Leu Leu  
35 40 45

Glu Lys Arg Val Gly Asp Arg Cys Cys Lys Gly Lys Arg Gly Cys Gly  
50 55 60

Arg Trp Cys Arg Asp His Ser Arg Cys Cys Gly Arg Arg  
65 70 75

<210> 36  
<211> 23  
<212> PRT  
<213> Conus bullatus

<220>  
<221> PEPTIDE  
<222> (1)..(23)  
<223> Xaa at residue 15 is Trp or bromo-Trp  
<400> 36  
Val Gly Asp Arg Cys Cys Lys Gly Lys Arg Gly Cys Gly Arg Xaa Cys  
1 5 10 15

Arg Asp His Ser Arg Cys Cys  
20

<210> 37  
<211> 331  
<212> DNA  
<213> Conus bullatus

<400> 37  
caagaaggat cgatagcagt tcatgatgtc taaactggga gtcttggtga ccactgtct 60  
gcttctgttt cccctttttg ctcttccgca ggatggagat caacctgcag accgacctgc 120  
agagcgtatg caggacgaca tttcatctga gcagaatccc ttgcttgaga agagagttgg 180  
tgaaagggtc tgcaaaaacg ggaagagggg gtgcggcaga tgggtgcagag atcactcacg 240  
ttgttgcggt cgacgataac gtgttgatga ccgaggcttt cgttatcacg gctacatcaa 300  
gtgtctagtg aataagtaaa acgattgcag t 331

<210> 38  
<211> 78  
<212> PRT  
<213> Conus bullatus

<400> 38  
Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
1 5 10 15

Pro Leu Phe Ala Leu Pro Gln Asp Gly Asp Gln Pro Ala Asp Arg Pro  
20 25 30

Ala Glu Arg Met Gln Asp Asp Ile Ser Ser Glu Gln Asn Pro Leu Leu  
35 40 45

Glu Lys Arg Val Gly Glu Arg Cys Cys Lys Asn Gly Lys Arg Gly Cys  
50 55 60

Gly Arg Trp Cys Arg Asp His Ser Arg Cys Cys Gly Arg Arg

12

65

70

75

&lt;210&gt; 39

&lt;211&gt; 24

&lt;212&gt; PRT

&lt;213&gt; Conus bullatus

&lt;220&gt;

&lt;221&gt; PEPTIDE

&lt;222&gt; (1)..(24)

&lt;223&gt; Xaa at residue 3 is Glu or gamma-carboxy Glu; Xaa at residue 16 is Trp or bromo-Tr

&lt;400&gt; 39

Val Gly Xaa Arg Cys Cys Lys Asn Gly Lys Arg Gly Cys Gly Arg Xaa  
1 5 10 15Cys Arg Asp His Ser Arg Cys Cys  
20

&lt;210&gt; 40

&lt;211&gt; 337

&lt;212&gt; DNA

&lt;213&gt; Conus bullatus

&lt;400&gt; 40

caagagggat cgaatagcagt tcattgatgtc taaactggga gtcttggtga ccatctgtct	60
gcttctgttt cccctttttg ctcttccgca ggacggagat caacctgcag accgacctgc	120
agagcgtatg caggacgacc ttcatctga gcagcatccc ttgtttgaga agagaattgt	180
tgacaggtgc tgcaacaaag ggaacgggaa gagggggtgc agcagatggt gcagagatca	240
ctcacgttgt tgcggtcgac gatgaactgt tgatgaccga ggctttggtt atcacggcta	300
catcaagtgt ctagtgaata agtaaacga ttgcagt	337

&lt;210&gt; 41

&lt;211&gt; 80

&lt;212&gt; PRT

&lt;213&gt; Conus bullatus

&lt;400&gt; 41

Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
1 5 10 15Pro Leu Phe Ala Leu Pro Gln Asp Gly Asp Gln Pro Ala Asp Arg Pro  
20 25 30Ala Glu Arg Met Gln Asp Asp Leu Ser Ser Glu Gln His Pro Leu Phe  
35 40 45Glu Lys Arg Ile Val Asp Arg Cys Cys Asn Lys Gly Asn Gly Lys Arg  
50 55 60Gly Cys Ser Arg Trp Cys Arg Asp His Ser Arg Cys Cys Gly Arg Arg  
65 70 75 80

&lt;210&gt; 42

&lt;211&gt; 26

&lt;212&gt; PRT

&lt;213&gt; Conus bullatus

&lt;220&gt;

13

&lt;221&gt; PEPTIDE

&lt;222&gt; (1)..(26)

&lt;223&gt; Xaa at residue 18 is Trp or bromo-Trp

&lt;400&gt; 42

Ile Val Asp Arg Cys Cys Asn Lys Gly Asn Gly Lys Arg Gly Cys Ser  
 1 5 10 15

Arg Xaa Cys Arg Asp His Ser Arg Cys Cys  
 20 25

&lt;210&gt; 43

&lt;211&gt; 337

&lt;212&gt; DNA

&lt;213&gt; Conus bullatus

&lt;400&gt; 43

caagaaggat cgatagcagt tcatgatgtc taaactggga gtcttggtga ccatctgtct 60  
 gcttctgttt cccctttttg ctcttccgca ggatggagat caacctgcag accgacctgc 120  
 tgagcgtatg caggacgaca ttcatctga gcggaatccc ttgtttgaga agagcgttgg 180  
 tttatattgc tgccgaccca aaccaacgg gcagatgatg tgcgacagat ggtgcgaaaa 240  
 aaactcacgt tgttgcggtc gacgataatg tgttgatgac cagctttgtt atcaaggcta 300  
 catcaagtat ctagtgaata agtaaacga ttgcagt 337

&lt;210&gt; 44

&lt;211&gt; 77

&lt;212&gt; PRT

&lt;213&gt; Conus bullatus

&lt;400&gt; 44

Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15

Pro Leu Phe Ala Leu Pro Gln Asp Gly Asp Gln Pro Ala Asp Arg Pro  
 20 25 30

Ala Glu Arg Met Gln Asp Asp Ile Ser Ser Asn Pro Leu Phe Glu Lys  
 35 40 45

Ser Val Gly Cys Cys Arg Pro Lys Pro Asn Gly Gln Met Met Cys Asp  
 50 55 60

Arg Trp Cys Glu Lys Asn Ser Arg Cys Cys Gly Arg Arg  
 65 70 75

&lt;210&gt; 45

&lt;211&gt; 27

&lt;212&gt; PRT

&lt;213&gt; Conus bullatus

&lt;220&gt;

&lt;221&gt; PEPTIDE

&lt;222&gt; (1)..(27)

<223> Xaa at residue 21 is Glu or gamma-carboxy Glu; Xaa at residue 8 and 10 is Pro or Hyp; Xaa at residue 19 is Trp or bromo-Trp; Xaa at residue 4 is Tyr, 125I-Tyr, mono-iodo-Tyr, di-iodo-Tyr, O-sulpho-Tyr or O-phospho-Ty

&lt;400&gt; 45

Val Gly Leu Xaa Cys Cys Arg Xaa Lys Xaa Asn Gly Gln Met Met Cys



14

1                    5                    10                    15

Asp Arg Xaa Cys Xaa Lys Asn Ser Arg Cys Cys  
                   20                    25

<210> 46  
 <211> 323  
 <212> DNA  
 <213> Conus bullatus  
 <400> 46  
 caagaaggat cgatagcagt tcatgatgtc taaactggga gttttgttga ccatctgtct 60  
 gcttctgttt ccccttactg ctcttccgat ggatggagat caatctgtag accgacctgc 120  
 agaacgtatg caggacgacc ttcatctga gcagcatccc ttgtttgttc agaaaagaag 180  
 gtgttgccgc gaaggcttga catgccccag atattggaaa aacagtcaga tttgtgcttg 240  
 ttgttaaatg acaacgtgtc gatgaccaac ttcggtatca cgactacgcc aagtgtctaa 300  
 tgaataagta aaacgattgc agt 323

<210> 47  
 <211> 74  
 <212> PRT  
 <213> Conus bullatus

<400> 47  
 Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1                    5                    10                    15

Pro Leu Thr Ala Leu Pro Met Asp Gly Asp Gln Ser Val Asp Arg Pro  
                   20                    25                    30

Ala Glu Arg Met Gln Asp Asp Leu Ser Ser Glu Gln His Pro Leu Phe  
                   35                    40                    45

Val Gln Lys Arg Arg Cys Cys Gly Glu Gly Leu Thr Cys Pro Arg Tyr  
                   50                    55                    60

Trp Lys Asn Ser Gln Ile Cys Ala Cys Cys  
 65                    70

<210> 48  
 <211> 22  
 <212> PRT  
 <213> Conus bullatus

<220>  
 <221> PEPTIDE  
 <222> (1)..(22)  
 <223> Xaa at residue 5 is Glu or gamma-carboxy Glu; Xaa at residue 10 is  
 s Pro or Hyp; Xaa at residue 13 is Trp or bromo-Trp; Xaa at resid  
 ue 12 is Tyr, 125I-Tyr, mono-iodo-Tyr, di-iodo-Tyr, O-sulpho-Tyr  
 or O-phospho-Ty

<400> 48  
 Arg Cys Cys Gly Xaa Gly Leu Thr Cys Xaa Arg Xaa Xaa Lys Asn Ser  
 1                    5                    10                    15

Gln Ile Cys Ala Cys Cys  
                   20

<210> 49  
 <211> 322

15

&lt;212&gt; DNA

<213> *Conus bullatus*

&lt;400&gt; 49

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caagagggat cgatagcagt tcatgatgtc taaactggga gtcttggtga ccatctgtct 60
gcttctgttt cccctttttg ctcttccgca ggatggagat caacctgcag accgacctgc 120

tgagcgtatg caggacgaca tttcatctga gcaggatccc ttgtttgttc agaaaagaag 180

gtgttgcggc gaaggcttga catgccccag atattggaaa aacagtcaga tttgtgcttg 240

ttgttaaatg acaacgtgtg atgaccaact tcggtatcac gactacgcca agtgtctaat 300

gaataagtaa aacgattgca gt 322

```

&lt;210&gt; 50

&lt;211&gt; 74

&lt;212&gt; PRT

<213> *Conus bullatus*

&lt;400&gt; 50

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Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Phe
1      5      10      15

Pro Leu Phe Ala Leu Pro Gln Asp Gly Asp Gln Pro Ala Asp Arg Pro
      20      25      30

Ala Glu Arg Met Gln Asp Asp Ile Ser Ser Glu Gln Asp Pro Leu Phe
      35      40      45

Val Gln Lys Arg Arg Cys Cys Gly Glu Gly Leu Thr Cys Pro Arg Tyr
      50      55      60

Trp Lys Asn Ser Gln Ile Cys Ala Cys Cys
65      70

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&lt;210&gt; 51

&lt;211&gt; 22

&lt;212&gt; PRT

<213> *Conus bullatus*

&lt;220&gt;

&lt;221&gt; PEPTIDE

&lt;222&gt; (1)..(22)

<223> Xaa at residue 5 is Glu or gamma-carboxy Glu; Xaa at residue 10 is Pro or Hyp; Xaa at residue 13 is Trp or bromo-Trp; Xaa at residue 12 is Tyr, 125I-Tyr, mono-iodo-Tyr, di-iodo-Tyr, O-sulpho-Tyr or O-phospho-Ty

&lt;400&gt; 51

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Arg Cys Cys Gly Xaa Gly Leu Thr Cys Xaa Arg Xaa Xaa Lys Asn Ser
1      5      10      15

Gln Ile Cys Ala Cys Cys
      20

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&lt;210&gt; 52

&lt;211&gt; 238

&lt;212&gt; DNA

<213> *Conus capitaneus*

&lt;400&gt; 52

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ggatccatga tgtctaaact gggagtcttg gtgaccatct gcctgcttct gtttccctt 60

getgcttttc cactggatgg aaatcaacct gcagaccacc ctgcaaagcg tacgcaagat 120

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gacagttcag ctgccctgat caatacctgg attgatcatt cccattcttg ctgcagggac 180

tgcggtgaag attgtgttgg ttgttgccgg taacgtgttg atgaccaact ttctcgag 238

<210> 53

<211> 70

<212> PRT

<213> Conus capitaneus

<400> 53

Gly Ser Met Met Ser Lys Leu Gly Val Leu Val Thr Ile Cys Leu Leu  
1 5 10 15

Leu Phe Pro Leu Ala Ala Phe Pro Leu Asp Gly Asn Gln Pro Ala Asp  
20 25 30

His Pro Ala Lys Arg Thr Gln Asp Asp Ser Ser Ala Ala Leu Ile Asn  
35 40 45

Thr Trp Ile Asp His Ser His Ser Cys Cys Arg Asp Cys Gly Glu Asp  
50 55 60

Cys Val Gly Cys Cys Arg  
65 70

<210> 54

<211> 15

<212> PRT

<213> Conus capitaneus

<220>

<221> PEPTIDE

<222> (1)..(15)

<223> Xaa at residue 8 is Glu or gamma-carboxy Glu

<400> 54

Ser Cys Cys Arg Asp Cys Gly Xaa Asp Cys Val Gly Cys Cys Arg  
1 5 10 15

<210> 55

<211> 323

<212> DNA

<213> Conus characteristicus

<400> 55

caagagggat cgatagcagt tcatgatgtc taaactggga gtcttggtga ccatctgtct 60

gcttctgttt ccccttactg ctcttccaat ggatggagat caacctgcag accaacctgc 120

agatcgtatg caggacgaca tttcatctga gcagtatccc ttgtttgata tgagaaaaag 180

gtgttgcggc cccggcggtt catgccccgt atatttcaga gacaatttta tttgtggttg 240

ttgttaaatg acaacgtgtc gatgaccaac ttcattatca cgactacgcc aagtgtctaa 300

tgaataagta aaatgattgc agt 323

<210> 56

<211> 74

<212> PRT

<213> Conus characteristicus

<400> 56

17

Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15  
 Pro Leu Thr Ala Leu Pro Met Asp Gly Asp Gln Pro Ala Asp Gln Pro  
 20 25 30  
 Ala Asp Arg Met Gln Asp Asp Ile Ser Ser Glu Gln Tyr Pro Leu Phe  
 35 40 45  
 Asp Met Arg Lys Arg Cys Cys Gly Pro Gly Gly Ser Cys Pro Val Tyr  
 50 55 60  
 Phe Arg Asp Asn Phe Ile Cys Gly Cys Cys  
 65 70

<210> 57  
 <211> 21  
 <212> PRT  
 <213> Conus characteristicus

<220>  
 <221> PEPTIDE  
 <222> (1)..(21)  
 <223> Xaa at residue 4 and 9 is Pro or Hyp; Xaa at residue 11 is Tyr, 1  
 25I-Tyr, mono-iodo-Tyr, di-iodo-Tyr, O-sulpho-Tyr or O-phospho-Ty

<400> 57  
 Cys Cys Gly Xaa Gly Gly Ser Cys Xaa Val Xaa Phe Arg Asp Asn Phe  
 1 5 10 15  
 Ile Cys Gly Cys Cys  
 20

<210> 58  
 <211> 316  
 <212> DNA  
 <213> Conus characteristicus

<400> 58  
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 cgagcgtatg caggacaaca tttcatctga gcagtatccc ttgtttgaga agagacgaga 180  
 ttgttgact cgcgcgaaga aatgcaaaga ccgacaatgc aaaccccgaga gatgttgccg 240  
 tggacgataa cgtgttgatg accaacttta tcacggctac gtcaagtgtt tagtgaataa 300  
 gtaaatgat tgcagt 316

<210> 59  
 <211> 75  
 <212> PRT  
 <213> Conus characteristicus

<400> 59  
 Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15  
 Pro Leu Thr Ala Leu Pro Met Asp Gly Asp Glu Pro Ala Asn Arg Pro  
 20 25 30  
 Val Glu Arg Met Gln Asp Asn Ile Ser Ser Glu Gln Tyr Pro Leu Phe



<212> PRT  
 <213> Conus characteristicus

<220>  
 <221> PEPTIDE  
 <222> (1)..(18)  
 <223> Xaa at residue 6 and 8 is Pro or Hyp; Xaa at residue 5 and 17 is Tyr, 125I-Tyr, mono-iodo-Tyr, di-iodo-Tyr, O-sulpho-Tyr or O-phospho-Ty

<400> 63  
 Arg Cys Cys Arg Xaa Xaa Cys Xaa Asp Ser Cys His Gly Ser Cys Cys  
 1 5 10 15

Xaa Lys

<210> 64  
 <211> 292  
 <212> DNA  
 <213> Conus characteristicus

<400> 64  
 caagagggat cgatagcagt tcatgatgtc taaactggga gccttggtga ccatctgtct 60  
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 acagcgtctg caggacogca ttccaactga agatcatccc ttatttgatc ccaacaaacg 180  
 gtgttgcccg ccggtggcat gcaacatggg atgcaagcct tgttggtggat gaccagcttt 240  
 gttatcgcgg tcttcatgaa gtgtcttaat gaataagtaa aatgattgca gt 292

<210> 65  
 <211> 69  
 <212> PRT  
 <213> Conus characteristicus

<400> 65  
 Met Met Ser Lys Leu Gly Ala Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15

Ser Leu Thr Ala Val Pro Leu Asp Gly Asp Gln His Ala Asp Gln Pro  
 20 25 30

Ala Gln Arg Leu Gln Asp Arg Ile Pro Thr Glu Asp His Pro Leu Phe  
 35 40 45

Asp Pro Asn Lys Arg Cys Cys Pro Pro Val Ala Cys Asn Met Gly Cys  
 50 55 60

Lys Pro Cys Cys Gly  
 65

<210> 66  
 <211> 15  
 <212> PRT  
 <213> Conus characteristicus

<220>  
 <221> PEPTIDE  
 <222> (1)..(15)  
 <223> Xaa at residue 3, 4 and 13 is Pro or Hyp

<400> 66  
 Cys Cys Xaa Xaa Val Ala Cys Asn Met Gly Cys Lys Xaa Cys Cys

20

1 5 10 15

<210> 67  
 <211> 293  
 <212> DNA  
 <213> Conus characteristicus

<400> 67  
 caagagggat cgatagcagt tcatgatgtc taaactggga gccttggtga ccatctgtct 60  
 acttctgttt tccctaactg ctgttccgct ggatggagat caacatgcag accaacctgc 120  
 agagcgtctg catgaccgcc ttccaactga aaatcatccc ttatatgata ccgtcaaacg 180  
 gtgttgcatg gattcggaat gcgactattc ttgctggcct tgctgtatgt ttggataacc 240  
 tttgttatcg cggcctcatc aagtgtctaa tgaataagta aaacgattgc agt 293

<210> 68  
 <211> 71  
 <212> PRT  
 <213> Conus characteristicus

<400> 68  
 Met Met Ser Lys Leu Gly Ala Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15  
 Ser Leu Thr Ala Val Pro Leu Asp Gly Asp Gln His Ala Asp Gln Pro  
 20 25 30  
 Ala Glu Arg Leu His Asp Arg Leu Pro Thr Glu Asn His Pro Leu Tyr  
 35 40 45  
 Asp Pro Val Lys Arg Cys Cys Asp Asp Ser Glu Cys Asp Tyr Ser Cys  
 50 55 60  
 Trp Pro Cys Cys Met Phe Gly  
 65 70

<210> 69  
 <211> 17  
 <212> PRT  
 <213> Conus characteristicus

<220>  
 <221> PEPTIDE  
 <222> (1)..(17)  
 <223> Xaa at residue 6 is Glu or gamma-carboxy Glu; Xaa at residue 13 is  
 s Pro or Hyp; Xaa at residue 12 is Trp or bromo-Trp; Xaa at resid  
 ue 9 is Tyr, 125I-Tyr, mono-iodo-Tyr, di-iodo-Tyr, O-sulpho-Tyr o  
 r O-phospho-Ty

<400> 69  
 Cys Cys Asp Asp Ser Xaa Cys Asp Xaa Ser Cys Xaa Xaa Cys Cys Met  
 1 5 10 15  
 Phe

<210> 70  
 <211> 232  
 <212> DNA  
 <213> Conus characteristicus

<400> 70  
 ggatccatga tgtctaaact gggagtcttg ttgaccatct gtctgttct gtttccctt 60

actgctgttc cgctggatgg agatcaacct gcagaccgac ctgcagagcg taagcaggac 120  
 gtttcatctg aacagcatcc cttctttgat cccgtcaaac ggtgttgccg ccggtgttac 180  
 atgggatgca tcccttggtg cttttaacgt gttgatgacc aactttctcg ag 232

<210> 71  
 <211> 68  
 <212> PRT  
 <213> Conus characteristicus

<400> 71  
 Gly Ser Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu  
 1 5 10 15  
 Leu Phe Pro Leu Thr Ala Val Pro Leu Asp Gly Asp Gln Pro Ala Asp  
 20 25 30  
 Arg Pro Ala Glu Arg Lys Gln Asp Val Ser Ser Glu Gln His Pro Phe  
 35 40 45  
 Phe Asp Pro Val Lys Arg Cys Cys Arg Arg Cys Tyr Met Gly Cys Ile  
 50 55 60

Pro Cys Cys Phe  
 65

<210> 72  
 <211> 14  
 <212> PRT  
 <213> Conus characteristicus

<220>  
 <221> PEPTIDE  
 <222> (1)..(14)  
 <223> Xaa at residue 11 is Pro or Hyp; Xaa at residue 6 is Tyr, 125I-Ty  
 r, mono-iodo-Tyr, di-iodo-Tyr, O-sulpho-Tyr or O-phospho-Ty

<400> 72  
 Cys Cys Arg Arg Cys Xaa Met Gly Cys Ile Xaa Cys Cys Phe  
 1 5 10

<210> 73  
 <211> 323  
 <212> DNA  
 <213> Conus circumcisis

<400> 73  
 caagaaggat cgatagcagt tcatgatgtc taaactgggg gtattgttga ccatctgtct 60  
 gcttctgttt ccccttactg ctcttccaat ggatggagat caacctgcag accaacctgc 120  
 agatcgtatg caggacgaca tttcatctga gcagtatccc ttgtttgata agagacgaaa 180  
 gtgttgcggc aaagacgggc catgccccaa atatttcaa gacaatttta tttgtggttg 240  
 ttgttaaatg acaacgtgtc gatgaccaac ttcgttatca cgattcgcca agtgtcttaa 300  
 tgaataagta aaatgattgc agt 323

<210> 74  
 <211> 74  
 <212> PRT  
 <213> Conus circumcisis



22

&lt;400&gt; 74

Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15

Pro Leu Thr Ala Leu Pro Met Asp Gly Asp Gln Pro Ala Asp Gln Pro  
 20 25 30

Ala Asp Arg Met Gln Asp Asp Ile Ser Ser Glu Gln Tyr Pro Leu Phe  
 35 40 45

Asp Lys Arg Arg Lys Cys Cys Gly Lys Asp Gly Pro Cys Pro Lys Tyr  
 50 55 60

Phe Lys Asp Asn Phe Ile Cys Gly Cys Cys  
 65 70

&lt;210&gt; 75

&lt;211&gt; 23

&lt;212&gt; PRT

&lt;213&gt; Conus circumciscus

&lt;220&gt;

&lt;221&gt; PEPTIDE

&lt;222&gt; (1)..(23)

<223> Xaa at residue 9 and 11 is Pro or Hyp; Xaa at residue 13 is Tyr,  
 125I-Tyr, mono-iodo-Tyr, di-iodo-Tyr, O-sulpho-Tyr or O-phospho-T  
 y

&lt;400&gt; 75

Arg Lys Cys Cys Gly Lys Asp Gly Xaa Cys Xaa Lys Xaa Phe Lys Asp  
 1 5 10 15

Asn Phe Ile Cys Gly Cys Cys  
 20

&lt;210&gt; 76

&lt;211&gt; 293

&lt;212&gt; DNA

&lt;213&gt; Conus dalli

&lt;400&gt; 76

caagagggat cgatagcagt tcatgatgtc taaactggga gccttggtga ccatctgtct 60

acttctgttt tcctaactg ctgttcgct ggatggagat caacatgcag accaacctgc 120

agagcgtctg caggaccgcc ttccaactga aaatcatccc ttatatgatc ccgtcaaacg 180

gtgttgcatg gattcggaat gcgactattc ttgctggcct tgctgtattt tatcataacc 240

tttgttatcg cggcctcatc aagtgtcaaa tgaataagta aaatgattgc agt 293

&lt;210&gt; 77

&lt;211&gt; 71

&lt;212&gt; PRT

&lt;213&gt; Conus dalli

&lt;400&gt; 77

Met Met Ser Lys Leu Gly Ala Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15

Ser Leu Thr Ala Val Pro Leu Asp Gly Asp Gln His Ala Asp Gln Pro  
 20 25 30

Ala Glu Arg Leu Gln Asp Arg Leu Pro Thr Glu Asn His Pro Leu Tyr  
 35 40 45

Asp Pro Val Lys Arg Cys Cys Asp Asp Ser Glu Cys Asp Tyr Ser Cys  
50 55 60

Trp Pro Cys Cys Ile Leu Ser  
65 70

<210> 78  
<211> 18  
<212> PRT  
<213> Conus dalli

<220>  
<221> PEPTIDE  
<222> (1)..(18)  
<223> Xaa at residue 6 is Glu or gamma-carboxy Glu; Xaa at residue 13 is Pro or Hyp; Xaa at residue 12 is Trp or bromo-Trp; Xaa at residue 9 is Tyr, 125I-Tyr, mono-iodo-Tyr, di-iodo-Tyr, O-sulpho-Tyr or O-phospho-Ty

<400> 78  
Cys Cys Asp Asp Ser Xaa Cys Asp Xaa Ser Cys Xaa Xaa Cys Cys Ile  
1 5 10 15

Leu Ser

<210> 79  
<211> 299  
<212> DNA  
<213> Conus dalli

<400> 79  
caagagggat cgatagcagt tcatgatgto taaactggga gtcttggtga ccatttgtct 60  
acttctgttt ccccttactg ctgttccact ggatggagat cagcctgcag accgacctgc 120  
agagcgtatg caggacggca tttcatctga acatcatcca ttttttgatt ccgtcaaaaa 180  
gaaacaacag tggtgcccgc cgggtggcatg caacatggga tgcgagcctt gttgtggatg 240  
accagctttg ttatcgccgc tcatgaagtg tcctaataaa taagtaaaac gattgcagt 299

<210> 80  
<211> 72  
<212> PRT  
<213> Conus dalli

<400> 80  
Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
1 5 10 15

Pro Leu Thr Ala Val Pro Leu Asp Gly Asp Gln Pro Ala Asp Arg Pro  
20 25 30

Ala Glu Arg Met Gln Asp Gly Ile Ser Ser Glu His His Pro Phe Phe  
35 40 45

Asp Ser Val Lys Lys Lys Gln Gln Cys Cys Pro Pro Val Ala Cys Asn  
50 55 60

Met Gly Cys Glu Pro Cys Cys Gly  
65 70

<210> 81  
<211> 17

<212> PRT  
<213> Conus dalli

<220>  
<221> PEPTIDE  
<222> (1)..(17)  
<223> Xaa at residue 1 is Gln or pyro-Glu; Xaa at residue 14 is Glu or gamma-carboxy Glu; Xaa at residue 5, 6 and 15 is Pro or Hyp

<400> 81  
Xaa Gln Cys Cys Xaa Xaa Val Ala Cys Asn Met Gly Cys Xaa Xaa Cys  
1 5 10 15

Cys

<210> 82  
<211> 290  
<212> DNA  
<213> Conus dalli

<400> 82  
caagaaggat cgatagcagt tcatgatgtc taaactggga gtcttggtga tcatatgtct 60  
atttctgttt ccccttactg ctgttcagct caatggagat cagcctgcag accaatctgc 120  
agagcgtatg caggacaaaa tttcatctga acatcatccc ttttttgatc ccgtcaaacg 180  
ttgttgcaac gcgggggttt gccgcttcgg atgcacgcct tgttggtggt gaccagcttt 240  
gttatcgcg cctcatcaag tgtctaata gaataagtaaaa tgattgcagt 290  
<210> 83  
<211> 69  
<212> PRT  
<213> Conus dalli

<400> 83  
Met Met Ser Lys Leu Gly Val Leu Leu Ile Ile Cys Leu Phe Leu Phe  
1 5 10 15  
Pro Leu Thr Ala Val Gln Leu Asn Gly Asp Gln Pro Ala Asp Gln Ser  
20 25 30  
Ala Glu Arg Met Gln Asp Lys Ile Ser Ser Glu His His Pro Phe Phe  
35 40 45  
Asp Pro Val Lys Arg Cys Cys Asn Ala Gly Phe Cys Arg Phe Gly Cys  
50 55 60

Thr Pro Cys Cys Trp  
65

<210> 84  
<211> 16  
<212> PRT  
<213> Conus dalli

<220>  
<221> PEPTIDE  
<222> (1)..(16)  
<223> Xaa at residue 13 is Pro or Hyp; Xaa at residue 16 is Trp or brom o-Tr

<400> 84  
Cys Cys Asn Ala Gly Phe Cys Arg Phe Gly Cys Thr Xaa Cys Cys Xaa  
1 5 10 15

25

<210> 85  
 <211> 288  
 <212> DNA  
 <213> Conus distans

<400> 85  
 caagagggat cgatagcagt tcatgatgtc taaactggga gtcttgctga coactcttct 60  
 gcttctgttt ccccttactg ctgttccgct ggatggagat caaccgcag acggacttgc 120  
 agagcgcagt caggacgaca gttcagctgc actgattaga gactggcttc ttcaaaccog 180  
 acagtgttgt gtgcatccat gcccatgcac gccttgctgt agatgaccag ctttgtcatc 240  
 gcggctacgt caagtatcta atgaataagt aagtaaaacg attgcagt 288

<210> 86  
 <211> 67  
 <212> PRT  
 <213> Conus distans

<400> 86  
 Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Phe Leu Leu Leu Phe  
 1 5 10 15  
 Pro Leu Thr Ala Val Pro Leu Asp Gly Asp Gln Pro Ala Asp Gly Leu  
 20 25 30  
 Ala Glu Arg Met Gln Asp Asp Ser Ser Ala Ala Leu Ile Arg Asp Trp  
 35 40 45  
 Leu Leu Gln Thr Arg Gln Cys Cys Val His Pro Cys Pro Cys Thr Pro  
 50 55 60  
 Cys Cys Arg  
 65

<210> 87  
 <211> 14  
 <212> PRT  
 <213> Conus distans

<220>  
 <221> PEPTIDE  
 <222> (1)..(14)  
 <223> Xaa at residue 1 is Gln or pyro-Glu; Xaa at residue 6, 8 and 11 i  
 s Pro or Hy

<400> 87  
 Xaa Cys Cys Val His Xaa Cys Xaa Cys Thr Xaa Cys Cys Arg  
 1 5 10

<210> 88  
 <211> 303  
 <212> DNA  
 <213> Conus ermineus

<400> 88  
 acctcaagag ggatcgatcg cagttcatga tgtctaaact gggagccttg ttgaccatct 60  
 gtctgcttct gtttccatt actgctcttc tgatggatgg agatcagcct gcagaccgac 120  
 ctgcagagcg tacggaggat gacatttcat ctgactacat tccctgttgc agttggccat 180

26

gcccccgata ctccaacggt aaacttggtt gtttttggtg ccttgatga taatgtgtg 240  
 atgaccaact ttgttatcac ggctacgtca agtgtctact gaataagtaa aatgattgca 300  
 gta 303

<210> 89  
 <211> 67  
 <212> PRT  
 <213> Conus ermineus

<400> 89  
 Met Met Ser Lys Leu Gly Ala Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15  
 Pro Ile Thr Ala Leu Leu Met Asp Gly Asp Gln Pro Ala Asp Arg Pro  
 20 25 30  
 Ala Glu Arg Thr Glu Asp Asp Ile Ser Ser Asp Tyr Ile Pro Cys Cys  
 35 40 45  
 Ser Trp Pro Cys Pro Arg Tyr Ser Asn Gly Lys Leu Val Cys Phe Cys  
 50 55 60  
 Cys Leu Gly  
 65

<210> 90  
 <211> 20  
 <212> PRT  
 <213> Conus ermineus

<220>  
 <221> PEPTIDE  
 <222> (1)..(20)  
 <223> Xaa at residue 5 and 7 is Pro or Hyp; Xaa at residue 4 is Trp or  
 bromo-Trp; Xaa at residue 9 is Tyr, 125I-Tyr, mono-iodo-Tyr, di-  
 iodo-Tyr, O-sulpho-Tyr or O-phospho-Ty

<400> 90  
 Cys Cys Ser Xaa Xaa Cys Xaa Arg Xaa Ser Asn Gly Lys Leu Val Cys  
 1 5 10 15  
 Phe Cys Cys Leu  
 20

<210> 91  
 <211> 241  
 <212> DNA  
 <213> Conus generalis

<400> 91  
 ggatccatga tgtctaaact gggagtcttg ttgaccatct gtctggttct gtttcccctt 60  
 actgtctttc cactggatgg agaacaacct gtagaccgac atgccgagca tatgcaggat 120  
 gacaattcag ctgcacagaa cccctgggtt attgccatca gacagtgttg cacgttctgc 180  
 aactttggat gccaaccttg ttgcctcacc tgataacgtg ttgatgacca actttctcga 240  
 g 241

<210> 92  
 <211> 70  
 <212> PRT

<213> *Conus generalis*

<400> 92

Gly Ser Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Val  
1 5 10 15

Leu Phe Pro Leu Thr Ala Leu Pro Leu Asp Gly Glu Gln Pro Val Asp  
20 25 30

Arg His Ala Glu His Met Gln Asp Asp Asn Ser Ala Ala Gln Asn Pro  
35 40 45

Trp Val Ile Ala Ile Arg Gln Cys Cys Thr Phe Cys Asn Phe Gly Cys  
50 55 60

Gln Pro Cys Cys Leu Thr  
65 70

<210> 93

<211> 16

<212> PRT

<213> *Conus generalis*

<220>

<221> PEPTIDE

<222> (1)..(16)

<223> Xaa at residue 1 is Gln or pyro-Glu; Xaa at residue 12 is Pro or  
Hy

<400> 93

Xaa Cys Cys Thr Phe Cys Asn Phe Gly Cys Gln Xaa Cys Cys Leu Thr  
1 5 10 15

<210> 94

<211> 241

<212> DNA

<213> *Conus generalis*

<400> 94

ggatccatga tgtctaaact gggagtcttg ttgaccatct gtctggttct gtttccctt 60

actgtcttc cactggatgg agaacaacct gtagaccgac atgccgagca tatgcaggat 120

gacaattcag ctgcacagaa cccctgggtt attgccatca gacagtgttg cacgttctgc 180

aactttggat gccagccttg ttgcgtcccc tgataacgtg ttgatgacca actttctcga 240

g 241

<210> 95

<211> 70

<212> PRT

<213> *Conus generalis*

<400> 95

Gly Ser Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Val  
1 5 10 15

Leu Phe Pro Leu Thr Ala Leu Pro Leu Asp Gly Glu Gln Pro Val Asp  
20 25 30

Arg His Ala Glu His Met Gln Asp Asp Asn Ser Ala Ala Gln Asn Pro  
35 40 45

Trp Val Ile Ala Ile Arg Gln Cys Cys Thr Phe Cys Asn Phe Gly Cys

50 55 60

Gln Pro Cys Cys Val Pro  
65 70

<210> 96  
<211> 16  
<212> PRT  
<213> Conus generalis

<220>  
<221> PEPTIDE  
<222> (1)..(16)  
<223> Xaa at residue 1 is Gln or pyro-Glu; Xaa at residue 12 and 16 is Pro or Hy

<400> 96  
Xaa Cys Cys Thr Phe Cys Asn Phe Gly Cys Gln Xaa Cys Cys Val Xaa  
1 5 10 15

<210> 97  
<211> 862  
<212> DNA  
<213> Conus geographus

<400> 97  
gtcgactcta gaggatccga caacaaagag tcaacccac tgccacgtca agagcgaagc 60  
gccacagcta agacaagagg gatcgatagc agttcatgat gtctaaactg ggagtcttgt 120  
tgaccatctg tctgcttctg tttcccotta ctgctcttcc gatggatgga gatgaacctg 180  
caaaccgacc tgtcgagcgt atgcaggaca acatttcato tgagcagtat cccttgtttg 240  
agaagagacg agattgttgc actccgccga agaaatgcaa agaccgacaa tgcaaaccac 300  
agagatgttg cgctggacga taacgtgttg atgaccaact ttatcacggc tacgtcaagt 360  
gttttagtga taagtaaaat gattgcagtc ttgctcagat ttgcttttgt gttttggtct 420  
aaagatcaat gaccaaaccg ttgttttgat gcggattgtc atatatttct cgattccaat 480  
ccaacactag atgatttaat cacgatagat taattttcta tcaatgcctt gatttttcgt 540  
ctgtcatatc agttttgttt atatttattt ttctgtcact gtctacacaa acgcatgcat 600  
gcacgcatgc acgcacacac gcacgcacgc tcgcacaaac atgcgcgcgc acgcacacac 660  
acacacacac acacaaacac acacacaagc aatcacacaa ttattgacat tatttattta 720  
ttcattgatg tatttgttat tcgtttgctt gtttttagaa tagtttgagg ccgtcttttt 780  
ggatttattt gaactgcttt attgtatacg agtacttcgt gctttgaaac actgctgaaa 840  
ataaaacaaa cactgacgta gc 862

<210> 98  
<211> 75  
<212> PRT  
<213> Conus geographus

<400> 98  
Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
1 5 10 15

## 29

Pro Leu Thr Ala Leu Pro Met Asp Gly Asp Glu Pro Ala Asn Arg Pro  
20 25 30

Val Glu Arg Met Gln Asp Asn Ile Ser Ser Glu Gln Tyr Pro Leu Phe  
35 40 45

Glu Lys Arg Arg Asp Cys Cys Thr Pro Pro Lys Lys Cys Lys Asp Arg  
50 55 60

Gln Cys Lys Pro Gln Arg Cys Cys Ala Gly Arg  
65 70 75

<210> 99

<211> 22

<212> PRT

<213> Conus geographus

<220>

<221> PEPTIDE

<222> (1)..(22)

<223> Xaa at residue 6, 7 and 17 is Pro or Hyp

<400> 99

Arg Asp Cys Cys Thr Xaa Xaa Lys Lys Cys Lys Asp Arg Gln Cys Lys  
1 5 10 15

Xaa Gln Arg Cys Cys Ala  
20

<210> 100

<211> 860

<212> DNA

<213> Conus geographus

<400> 100

ggccagacga caacaaagag tcaacccac tgccacgtca agagcgaagc gccacagcta 60  
agacaagagg gatcgatagc agttcatgat gtctaaactg ggagtcttgt tgaccatctg 120  
tctgcttctg tttcccctta ctgctcttcc gatggatgga gatgaacctg caaaccgacc 180  
tgctgagcgt atgcaggaca acatttcata tgagcagtat cccttgtttg agaagagacg 240  
agattgttgc actccgccga ggaaatgcaa agaccgacga tgcaaaccga tgaaatgttg 300  
cgctggacga taacgtgttg atgaccaact ttatcacggc tagctcagtg tttagtgaat 360  
aagtaaaatg attgcagtct tgctcagatt gcttttgtgt tttggtctaa gatcaatgac 420  
caaaccttg ttttgatgcg gattgtcata ttttctcga ttccaatcca aactagatg 480  
atttaacac gatagattaa ttttctatca atgccttgat tttcgtctg tcatatcagt 540  
tttgtttata tttatttttt cgtcactgtc tacacaaacg catgcatgca cgcattgcacg 600  
cacacacgca cgcacgctcg cacaacatg cgcgcgcacg cacacacaca cacacacaca 660  
aacacacaca cgaagcaatc acacaattag ttgacattat ttatttattc attgatgtat 720  
ttgttattcg tttgcttgtt tttagaatag tttagggcgg tcttttttga tttatttgaa 780  
ctgctttatt gtatacagat acttcgtgct ttgaaacact gctgaaaata aaacaaacac 840  
tgacgtagca aaaaaaaaaa 860



30

<210> 101  
 <211> 75  
 <212> PRT  
 <213> Conus geographus

<400> 101  
 Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15  
 Pro Leu Thr Ala Leu Pro Met Asp Gly Asp Glu Pro Ala Asn Arg Pro  
 20 25 30  
 Val Glu Arg Met Gln Asp Asn Ile Ser Ser Glu Gln Tyr Pro Leu Phe  
 35 40 45  
 Glu Lys Arg Arg Asp Cys Cys Thr Pro Pro Arg Lys Cys Lys Asp Arg  
 50 55 60  
 Arg Cys Lys Pro Met Lys Cys Cys Ala Gly Arg  
 65 70 75

<210> 102  
 <211> 22  
 <212> PRT  
 <213> Conus geographus

<220>  
 <221> PEPTIDE  
 <222> (1)..(22)  
 <223> Xaa at residue 6, 7 and 17 is Pro or Hyp

<400> 102  
 Arg Asp Cys Cys Thr Xaa Xaa Arg Lys Cys Lys Asp Arg Arg Cys Lys  
 1 5 10 15  
 Xaa Met Lys Cys Cys Ala  
 20

<210> 103  
 <211> 22  
 <212> PRT  
 <213> Conus geographus

<220>  
 <221> PEPTIDE  
 <222> (1)..(22)  
 <223> Xaa at residue 6, 7 and 17 is Pro or Hyp

<400> 103  
 Arg Asp Cys Cys Thr Xaa Xaa Lys Lys Cys Lys Asp Arg Arg Cys Lys  
 1 5 10 15  
 Xaa Leu Lys Cys Cys Ala  
 20

<210> 104  
 <211> 321  
 <212> DNA  
 <213> Conus gloriamaris

<400> 104  
 ctactatag gaattcgagc tcggtacacg ggatcgatag cagttcatga tgtctaaact 60  
 gggagccttg ttgaccatct gtctacttct gttttcccta actgctgttc cgctggatgg 120

agatcaacat gcagaccaac ctgcagagcg tctgcatgac cgccttccaa ctgaaaatca 180  
 tcccttatat gatcccgta aacggtgttg cgatgattcg gaatgcgact attcttgctg 240  
 gccttgctgt atgtttggat aacctttgtt atcgcggcct cgataagtgt ctaatgaata 300  
 agtaaaacga ttgcagtagg c 321

<210> 105  
 <211> 71  
 <212> PRT  
 <213> Conus gloriamaris

<400> 105  
 Met Met Ser Lys Leu Gly Ala Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15  
 Ser Leu Thr Ala Val Pro Leu Asp Gly Asp Gln His Ala Asp Gln Pro  
 20 25 30  
 Ala Glu Arg Leu His Asp Arg Leu Pro Thr Glu Asn His Pro Leu Tyr  
 35 40 45  
 Asp Pro Val Lys Arg Cys Cys Asp Asp Ser Glu Cys Asp Tyr Ser Cys  
 50 55 60  
 Trp Pro Cys Cys Met Phe Gly  
 65 70

<210> 106  
 <211> 17  
 <212> PRT  
 <213> Conus gloriamaris

<220>  
 <221> PEPTIDE  
 <222> (1)..(17)  
 <223> Xaa at residue is 6 Glu or gamma-carboxy Glu; Xaa at residue 13 is  
 s Pro or Hyp; Xaa at residue 12 is Trp or bromo-Trp; Xaa at resid  
 ue 9 is Tyr, 125I-Tyr, mono-iodo-Tyr, di-iodo-Tyr, O-sulpho-Tyr o  
 r O-phospho-Ty

<400> 106  
 Cys Cys Asp Asp Ser Xaa Cys Asp Xaa Ser Cys Xaa Xaa Cys Cys Met  
 1 5 10 15

Phe

<210> 107  
 <211> 257  
 <212> DNA  
 <213> Conus gloriamaris

<400> 107  
 gttcatgatg tctaaactgg gagtcttgtt gatcatctgt ctacttctgt ttccccttac 60  
 tgctgttccg ctggatggag atcaacctgc agaccgatat gcagagcgta tgcaggacga 120  
 catttcatct gaacatcatc ccatgtttga tgccgtcaga ggggtgttgcc atctgttggc 180  
 atgccgcttc ggatgctcgc cttgttgttg gtgatcagct ttgttatcgc ggcctcatca 240  
 agtgactcta atgcaaa 257  
 <210> 108  
 <211> 69

32

&lt;212&gt; PRT

&lt;213&gt; Conus gloriamaris

&lt;400&gt; 108

Met Met Ser Lys Leu Gly Val Leu Leu Ile Ile Cys Leu Leu Leu Phe  
 1 5 10 15

Pro Leu Thr Ala Val Pro Leu Asp Gly Asp Gln Pro Ala Asp Arg Tyr  
 20 25 30

Ala Glu Arg Met Gln Asp Asp Ile Ser Ser Glu His His Pro Met Phe  
 35 40 45

Asp Ala Val Arg Gly Cys Cys His Leu Leu Ala Cys Arg Phe Gly Cys  
 50 55 60

Ser Pro Cys Cys Trp  
 65

&lt;210&gt; 109

&lt;211&gt; 17

&lt;212&gt; PRT

&lt;213&gt; Conus gloriamaris

&lt;220&gt;

&lt;221&gt; PEPTIDE

&lt;222&gt; (1)..(17)

&lt;223&gt; Xaa at residue 14 is Pro or Hyp; Xaa at residue 17 is Trp or brom o-Tr

&lt;400&gt; 109

Gly Cys Cys His Leu Leu Ala Cys Arg Phe Gly Cys Ser Xaa Cys Cys  
 1 5 10 15

Xaa

&lt;210&gt; 110

&lt;211&gt; 471

&lt;212&gt; DNA

&lt;213&gt; Conus gloriamaris

&lt;400&gt; 110

gagacgacaa ggaacagtca accccacagc cacgccaaga gcagacagcc acagctacgt 60  
 gaagaagggt ggagagaggt tcgtgatgtt gaaaatggga gtggtgctat tcatcttcct 120  
 ggtactgttt cccctggcaa cgctccagct ggatgcagat caacctgtag aacgatatgc 180  
 ggagaacaaa cagctcctca acccagatga aaggaggga atcatattgc atgctctggg 240  
 gacgcgatgc tgttcttggg atgtgtgcga ccacccgagt tgtacttgct gcggcggtta 300  
 gcgccgaaca tccatggcgc tgtgctgggc ggttttatcc aacaacgaca gogtttgttg 360  
 atttcatgta tcattgcgcc cacgtctctt gtctaagaat gacgaacatg attgcactct 420  
 ggttcagatt tcgtgttctt ttctgacaat aaatgacaaa actccaaaaa a 471

&lt;210&gt; 111

&lt;211&gt; 71

&lt;212&gt; PRT

&lt;213&gt; Conus gloriamaris

&lt;400&gt; 111

Met Leu Lys Met Gly Val Val Leu Phe Ile Phe Leu Val Leu Phe Pro

33

1                    5                    10                    15  
 Leu Ala Thr Leu Gln Leu Asp Ala Asp Gln Pro Val Glu Arg Tyr Ala  
                   20                    25                    30  
 Glu Asn Lys Gln Leu Leu Asn Pro Asp Glu Arg Arg Glu Ile Ile Leu  
                   35                    40                    45  
 His Ala Leu Gly Thr Arg Cys Cys Ser Trp Asp Val Cys Asp His Pro  
                   50                    55                    60  
 Ser Cys Thr Cys Cys Gly Gly  
 65                    70

<210> 112  
 <211> 16  
 <212> PRT  
 <213> Conus gloriamaris  
 <220>  
 <221> PEPTIDE  
 <222> (1)..(16)  
 <223> Xaa at residue 10 is Pro or Hyp; Xaa at residue 4 is Trp or bromo  
 -Tr

<400> 112  
 Cys Cys Ser Xaa Asp Val Cys Asp His Xaa Ser Cys Thr Cys Cys Gly  
 1                    5                    10                    15

<210> 113  
 <211> 304  
 <212> DNA  
 <213> Conus laterculatus

<400> 113  
 cgacctcaag aaggatcgat agcagttcat gatgtctaaa ctgggagtct tgttgaccat 60  
 ctgtctgctt ctgtttcccc ttactgctct tccgatggat ggagatcaac ctgcagaccg 120  
 acctgcagag cgtatgcagg acgtttcatc tgaacagcat cccttgatg atcccgtaa 180  
 acggtgttgc gactggccat gcagcggatg catcccttgt tgctaatagt aacaacgtgt 240  
 tgataaccaa ctttcttacc acgactacgt caagtgtcta atgaataagt aaaatgattg 300  
 cagt 304

<210> 114  
 <211> 65  
 <212> PRT  
 <213> Conus laterculatus

<400> 114  
 Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1                    5                    10                    15  
 Pro Leu Thr Ala Leu Pro Met Asp Gly Asp Gln Pro Ala Asp Arg Pro  
                   20                    25                    30  
 Ala Glu Arg Met Gln Asp Val Ser Ser Glu Gln His Pro Leu Tyr Asp  
                   35                    40                    45  
 Pro Val Lys Arg Cys Cys Asp Trp Pro Cys Ser Gly Cys Ile Pro Cys  
                   50                    55                    60

Cys

65

<210> 115  
 <211> 13  
 <212> PRT  
 <213> Conus laterculatus

<220>  
 <221> PEPTIDE  
 <222> (1)..(13)  
 <223> Xaa at residue 5 and 11 is Pro or Hyp; Xaa at residue 4 is Trp or bromo-Trp

<400> 115  
 Cys Cys Asp Xaa Xaa Cys Ser Gly Cys Ile Xaa Cys Cys  
 1 5 10

<210> 116  
 <211> 313  
 <212> DNA  
 <213> Conus laterculatus

<400> 116  
 cgacctcaag aaggatogat agcagttcat gatgtctaaa ctgggagtct tggtgaccat 60  
 ctgtctgctt ctgtttcccc ttactgctct ggatggagat caacctgcag accgacttgc 120  
 agagcgtatg caggacgaca tttcatctga gcagcatccc ttgaaaaga gacgagactg 180  
 ttgcacacct ccgaagaaat gcagagaccg acaatgcaaa cctgcacggt gttgcggagg 240  
 ataacgtgtt gatgaccaac tttgttatca cggctacgtc aagtgtctag tgaataagta 300  
 aaacgattgc agt 313

<210> 117  
 <211> 71  
 <212> PRT  
 <213> Conus laterculatus

<400> 117  
 Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15

Pro Leu Thr Ala Leu Asp Gly Asp Gln Pro Ala Asp Arg Leu Ala Glu  
 20 25 30

Arg Met Gln Asp Asp Ile Ser Ser Glu Gln His Pro Phe Glu Lys Arg  
 35 40 45

Arg Asp Cys Cys Thr Pro Pro Lys Lys Cys Arg Asp Arg Gln Cys Lys  
 50 55 60

Pro Ala Arg Cys Cys Gly Gly  
 65 70

<210> 118  
 <211> 22  
 <212> PRT  
 <213> Conus laterculatus

<220>  
 <221> PEPTIDE  
 <222> (1)..(22)  
 <223> Xaa at residue 6, 17 and 17 is Pro or Hyp

&lt;400&gt; 118

Arg Asp Cys Cys Thr Xaa Xaa Lys Lys Cys Arg Asp Arg Gln Cys Lys  
 1 5 10 15

Xaa Ala Arg Cys Cys Gly  
 20

&lt;210&gt; 119

&lt;211&gt; 314

&lt;212&gt; DNA

&lt;213&gt; Conus laterculatus

&lt;400&gt; 119

gggategata gcagttcatg atgtctaaac tgggagtctt gttgaccatc tgtctgcttc 60  
 tgtttccctt tactgtctctt ccgatggatg gagatcaact tgcaagccga tctgcagagc 120  
 gtatgcagga caacatttca tctgagcagc atcacctctt tgaaaagaga cgaccacat 180  
 gttgcaccta tgacgggagt tgcctaaaag aatcatgcat gcgtaaagct tgttgccgat 240  
 gataacgtgt tgatgaccaa ctttgttctc acggctactc aagtgtctaa tgaataagta 300  
 aaatgattgc agta 314

&lt;210&gt; 120

&lt;211&gt; 74

&lt;212&gt; PRT

&lt;213&gt; Conus laterculatus

&lt;400&gt; 120

Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15

Pro Leu Thr Ala Leu Pro Met Asp Gly Asp Gln Leu Ala Arg Arg Ser  
 20 25 30

Ala Glu Arg Met Gln Asp Asn Ile Ser Ser Glu Gln His His Leu Phe  
 35 40 45

Glu Lys Arg Arg Pro Pro Cys Cys Thr Tyr Asp Gly Ser Cys Leu Lys  
 50 55 60

Glu Ser Cys Met Arg Lys Ala Cys Cys Gly  
 65 70

&lt;210&gt; 121

&lt;211&gt; 22

&lt;212&gt; PRT

&lt;213&gt; Conus laterculatus

&lt;220&gt;

&lt;221&gt; PEPTIDE

&lt;222&gt; (1)..(22)

<223> Xaa at residue 14 is Glu or gamma-carboxy Glu; Xaa at residue 2 and 3 is Pro or Hyp; Xaa at residue 7 is Tyr, 125I-Tyr, mono-iodo-Tyr, di-iodo-Tyr, O-sulpho-Tyr or O-phospho-Ty

&lt;400&gt; 121

Arg Xaa Xaa Cys Cys Thr Xaa Asp Gly Ser Cys Leu Lys Xaa Ser Cys  
 1 5 10 15

Met Arg Lys Ala Cys Cys  
 20

<210> 122  
 <211> 314  
 <212> DNA  
 <213> Conus laterculatus

<400> 122  
 gggatcgata gcagttcatg atgtctaaac tgggagtctt gttgaccacc tgtctgcttc 60  
 tgtttccctt tactgctctt ccgatggatg gagatcaact tgcacgccga cctgcagagc 120  
 gtatgcagga caacatttca tctgagcagc atcccttctt tgaaaggaga cgaccacat 180  
 gttgcaccta tgacggggagt tgcctaaaag aatcatgcaa gcgtaaagct tgttgccgat 240  
 aataacgtgt tgatgaccaa ctttgttatc acggctactc aagtgtctaa tgaataagta 300  
 aaatgattgc agta 314

<210> 123  
 <211> 74  
 <212> PRT  
 <213> Conus laterculatus

<400> 123  
 Met Met Ser Lys Leu Gly Val Leu Leu Thr Thr Cys Leu Leu Leu Phe  
 1 5 10 15  
 Pro Leu Thr Ala Leu Pro Met Asp Gly Asp Gln Leu Ala Arg Arg Pro  
 20 25 30  
 Ala Glu Arg Met Gln Asp Asn Ile Ser Ser Glu Gln His Pro Phe Phe  
 35 40 45  
 Glu Arg Arg Arg Pro Pro Cys Cys Thr Tyr Asp Gly Ser Cys Leu Lys  
 50 55 60  
 Glu Ser Cys Lys Arg Lys Ala Cys Cys Gly  
 65 70

<210> 124  
 <211> 22  
 <212> PRT  
 <213> Conus laterculatus

<220>  
 <221> PEPTIDE  
 <222> (1)..(22)  
 <223> Xaa at residue 14 is Glu or gamma-carboxy Glu; Xaa at residue 2 and 3 is Pro or Hyp; Xaa at residue 7 is Tyr, 125I-Tyr, mono-iodo-Tyr, di-iodo-Tyr, O-sulpho-Tyr or O-phospho-Ty

<400> 124  
 Arg Xaa Xaa Cys Cys Thr Xaa Asp Gly Ser Cys Leu Lys Xaa Ser Cys  
 1 5 10 15  
 Lys Arg Lys Ala Cys Cys  
 20

<210> 125  
 <211> 247  
 <212> DNA  
 <213> Conus leopardus

<400> 125

37

ggatccatga tgtctaaact gggagtcttg ttgaccgtct gtctgcttct gtttcccctt 60  
 actgctcttc ggctgggttg agatcaacct gcagagcgac ctgcaaagcg tacgcaggac 120  
 gacattccag atggacagca tccgttaaat gataggcaga taaactgttg cccgtggcca 180  
 tgccctagta catgcgcgca tcaatgctgc cattaatgat aacgtgttga tgaccaactt 240  
 tctcgag 247

<210> 126  
 <211> 71  
 <212> PRT  
 <213> Conus leopardus

<400> 126  
 Gly Ser Met Met Ser Lys Leu Gly Val Leu Leu Thr Val Cys Leu Leu  
 1 5 10 15  
 Leu Phe Pro Leu Thr Ala Leu Arg Leu Val Gly Asp Gln Pro Ala Glu  
 20 25 30  
 Arg Pro Ala Lys Arg Thr Gln Asp Asp Ile Pro Asp Gly Gln His Pro  
 35 40 45  
 Leu Asn Asp Arg Gln Ile Asn Cys Cys Pro Trp Pro Cys Pro Ser Thr  
 50 55 60  
 Cys Arg His Gln Cys Cys His  
 65 70

<210> 127  
 <211> 19  
 <212> PRT  
 <213> Conus leopardus

<220>  
 <221> PEPTIDE  
 <222> (1)..(19)  
 <223> Xaa at residue 1 is Gln or pyro-Glu; Xaa at residue 6, 8 and 10 is Pro or Hyp; Xaa at residue 7 is Trp or bromo-Tr

<400> 127  
 Xaa Ile Asn Cys Cys Xaa Xaa Xaa Cys Xaa Ser Thr Cys Arg His Gln  
 1 5 10 15  
 Cys Cys His

<210> 128  
 <211> 244  
 <212> DNA  
 <213> Conus lividus

<400> 128  
 ggatccatga tgtctaaact gggagtcttg ttgaccgtct gtctgcttct gtttcccctt 60  
 actgctcttc ggctgggttag agatcaacct gcagagcgac ctgcaaagcg tacgcaggac 120  
 gacattccaa atggacagga tccgttaatt gataggcaga taaattgttg cccttgcca 180  
 tgccctgatt catgccacta tcaatgctgc cactgataac gtgttgatga ccaactttct 240  
 cgag 244

<210> 129



38

<211> 71  
 <212> PRT  
 <213> Conus lividus

<400> 129  
 Gly Ser Met Met Ser Lys Leu Gly Val Leu Leu Thr Val Cys Leu Leu  
 1 5 10 15  
 Leu Phe Pro Leu Thr Ala Leu Arg Leu Val Arg Asp Gln Pro Ala Glu  
 20 25 30  
 Arg Pro Ala Lys Arg Thr Gln Asp Asp Ile Pro Asn Gly Gln Asp Pro  
 35 40 45  
 Leu Ile Asp Arg Gln Ile Asn Cys Cys Pro Trp Pro Cys Pro Asp Ser  
 50 55 60  
 Cys His Tyr Gln Cys Cys His  
 65 70

<210> 130  
 <211> 19  
 <212> PRT  
 <213> Conus lividus

<220>  
 <221> PEPTIDE  
 <222> (1)..(19)  
 <223> Xaa at residue 1 is Gln or pyro-Glu; Xaa at residue 6, 8 and 10 is Pro or Hyp; Xaa at residue 7 is Trp or bromo-Trp; Xaa at residue 15 is Tyr, 125I-Tyr, mono-iodo-Tyr, di-iodo-Tyr, O-sulpho-Tyr or O-phospho-Ty

<400> 130  
 Xaa Ile Asn Cys Cys Xaa Xaa Xaa Cys Xaa Asp Ser Cys His Xaa Gln  
 1 5 10 15

Cys Cys His

<210> 131  
 <211> 275  
 <212> DNA  
 <213> Conus lynceus

<400> 131  
 aaggatcgat agcagttcat gatgtctaaa ctgggagtct tgttgaccat ctgtctgctt 60  
 ctgtttcccc ttactgctct tccgatggat ggagatcaat ctgcagaccg acttgcagag 120  
 cgtatgcagg acaacatttc atctgagcag catcccttct ttgaaaagag aggacgagac 180  
 tgttgcacac ctccgaggaa atgcagagac cgagcctgca aacctcaacg ttgttgcgga 240  
 ggataagctg ttgatgacca actttgttat acggc 275

<210> 132  
 <211> 75  
 <212> PRT  
 <213> Conus lynceus

<400> 132  
 Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Phe  
 1 5 10 15

Pro Leu Thr Ala Leu Pro Met Asp Gly Asp Gln Ser Ala Asp Arg Leu

39

20 25 30

Ala Glu Arg Met Gln Asp Asn Ile Ser Ser Glu Gln His Pro Phe Phe  
35 40 45

Glu Lys Arg Gly Arg Asp Cys Cys Thr Pro Pro Arg Lys Cys Arg Asp  
50 55 60

Arg Ala Cys Lys Pro Gln Arg Cys Cys Gly Gly  
65 70 75

<210> 133  
<211> 23  
<212> PRT  
<213> Conus lynceus

<220>  
<221> PEPTIDE  
<222> (1)..(23)  
<223> Xaa at residue 7, 8 and 18 is Pro or Hyp

<400> 133  
Gly Arg Asp Cys Cys Thr Xaa Xaa Arg Lys Cys Arg Asp Arg Ala Cys  
1 5 10 15

Lys Xaa Gln Arg Cys Cys Gly  
20

<210> 134  
<211> 803  
<212> DNA  
<213> Conus magus

<400> 134  
caagagggat cgatagcagt tcatgatgtc taaactggga gtcttggtga ccatctgtct 60  
gcttctgttt ccccttactg ctcttccgat gtagggagat gaacctgcaa accgacctgt 120  
cgagcgtatg caggacaaca tttcatctga gcagtatccc ttgtttgaga agagacgaga 180  
ttgttgact cgcgcaaga aatgcaaaga ccgacaatgc aaacccaga gatgttgccg 240  
tggaagataa cgtgttgatg accaacttta tcacggctac gtcaagtgtt tagtgaataa 300  
gtaaaatgat tgcagtcttg ctcagatttg cttttgtgtt ttggtctaaa gatcaatgac 360  
caaaccgttg ttttgatgcg gattgtcata ttttctcga ttccaatcca aactagatg 420  
atttaatcac gatagattaa ttttctatca atgccttgat ttttcgtctg tcatatcagt 480  
tttgtttata tttatttttt cgtcactgtc tacacaaacg catgcatgca cgcattgcacg 540  
cacacacgca cgcacgctcg cacaacatg cgcgcgcacg cacacacaca cacacacaca 600  
caaacacaca caggaagcaa tcacacaatt agttgacatt atttatttat tcattgatgt 660  
atttgattat cgtttgcttg tttttagaat agtttgaggc cgtctttttg gatttatttg 720  
aactgcttta ttgtatacga gtacttcgtg cggggaaaca ctgctgaaaa taaaacaaac 780  
actgacgtag caaaaaaaaaaaa aaa 803

<210> 135  
<211> 75  
<212> PRT

40

&lt;213&gt; Conus magus

&lt;400&gt; 135

Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
1 5 10 15Pro Leu Thr Ala Leu Pro Met Asp Gly Asp Glu Pro Ala Asn Arg Pro  
20 25 30Val Glu Arg Met Gln Asp Asn Ile Ser Ser Glu Gln Tyr Pro Leu Phe  
35 40 45Glu Lys Arg Arg Asp Cys Cys Thr Pro Pro Lys Lys Cys Lys Asp Arg  
50 55 60Gln Cys Lys Pro Gln Arg Cys Cys Ala Gly Arg  
65 70 75

&lt;210&gt; 136

&lt;211&gt; 22

&lt;212&gt; PRT

&lt;213&gt; Conus magus

&lt;220&gt;

&lt;221&gt; PEPTIDE

&lt;222&gt; (1)..(22)

&lt;223&gt; Xaa at residue 6 and 7 is Pro or Hyp

&lt;400&gt; 136

Arg Asp Cys Cys Thr Xaa Xaa Lys Lys Cys Lys Asp Arg Gln Cys Lys  
1 5 10 15Xaa Gln Arg Cys Cys Ala  
20

&lt;210&gt; 137

&lt;211&gt; 656

&lt;212&gt; DNA

&lt;213&gt; Conus magus

&lt;400&gt; 137

caagagggat cgatagcagt tcatgatgtc taaactggga gtcttgttga ccatctgtct 60

gcttctgttt ccccttactg ctcttccaat ggatggagat caacctgcag accaacctgc 120

agatcgtatg caggacgaca tttcatctga gcagtatccc ttgtttgata tgagaaaaag 180

gtgttgccgc cccggcgggt catgccccgt atatttcaga gacaatttta tttgtggttg 240

ttgttaaatg acaacgtgtc gatgaccaac ttcattatca cgactacgcc aagtgtctaa 300

tgaataaata aaatgattgc agtctcgctc agatttgctt ttgtattttg gtctaaagat 360

caatgaccaa accgttgttt tgggtgtggat tttcatatat ttctcgagtc ctatccaaca 420

ctagatgatt taatcacgat agatctgatt tttttatcaa aggcttggtt tttcgtctgt 480

cacatcagtt ttgtttatat ttaatttttc gtcactgatt acacacacgc atgaacgcac 540

agagtactaa cacatacaca cacacacaca cacacacaca cacacacaca cacacacaca 600

cacacacaca caccgcgcgc cgcggcgcga tctagtagcg ccgcgcgcac acacac 656

&lt;210&gt; 138

&lt;211&gt; 74

&lt;212&gt; PRT

41

&lt;213&gt; Conus magus

&lt;400&gt; 138

Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Phe  
1 5 10 15Pro Leu Thr Ala Leu Pro Met Asp Gly Asp Gln Pro Ala Asp Gln Pro  
20 25 30Ala Asp Arg Met Gln Asp Asp Ile Ser Ser Glu Gln Tyr Pro Leu Phe  
35 40 45Asp Met Arg Lys Arg Cys Cys Gly Pro Gly Gly Ser Cys Pro Val Tyr  
50 55 60Phe Arg Asp Asn Phe Ile Cys Gly Cys Cys  
65 70

&lt;210&gt; 139

&lt;211&gt; 21

&lt;212&gt; PRT

&lt;213&gt; Conus magus

&lt;220&gt;

&lt;221&gt; PEPTIDE

&lt;222&gt; (1)..(21)

<223> Xaa at residue 4 and 9 is Pro or Hyp; Xaa at residue is 11 Tyr, 1  
25I-Tyr, mono-iodo-Tyr, di-iodo-Tyr, O-sulpho-Tyr or O-phospho-Ty

&lt;400&gt; 139

Cys Cys Gly Xaa Gly Gly Ser Cys Xaa Val Xaa Phe Arg Asp Asn Phe  
1 5 10 15Ile Cys Gly Cys Cys  
20

&lt;210&gt; 140

&lt;211&gt; 594

&lt;212&gt; DNA

&lt;213&gt; Conus magus

&lt;400&gt; 140

caagagggat cgatagcagt tcatgatgtc taaactggga gtcttggtga ccactctgttt 60

gcttctgttt ccccttactg ctcttccgag ggatggagat caatctgtag accgacctgc 120

agagcgtatg caggacgaca ttcatctga gctgcatccc ttgtcaatca gaaaaagaat 180

gtgttgccgc gagagtgcgc catgccccag ctatttcaga aacagtcaga tttgtcattg 240

ttgttaaatg acaacgtgtc gatgaccacc ttggttatca cgactaatga taagtaaaat 300

gattgcagtc tcgctcagat ttgcttttgt attttggct aaagatcaat gaccaaaccg 360

ttgttttgat gtggattttc atatatttct cgagtcctat ccaacactag atgatttaat 420

cacgatagat ctgatttttt tatcaaagcc ttgggttttc gtctgtcaca tcagttttgt 480

ttatatattaa tttttcgtca ctgattacac acacgcatga acgcacagac gtactaacac 540

atacacacac acacacacac acaacacac acacacacac acacacacac acac 594

&lt;210&gt; 141

&lt;211&gt; 74

42

&lt;212&gt; PRT

&lt;213&gt; Conus magus

&lt;400&gt; 141

Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15

Pro Leu Thr Ala Leu Pro Arg Asp Gly Asp Gln Ser Val Asp Arg Pro  
 20 25 30

Ala Glu Arg Met Gln Asp Asp Ile Ser Ser Glu Leu His Pro Leu Ser  
 35 40 45

Ile Arg Lys Arg Met Cys Cys Gly Glu Ser Ala Pro Cys Pro Ser Tyr  
 50 55 60

Phe Arg Asn Ser Gln Ile Cys His Cys Cys  
 65 70

&lt;210&gt; 142

&lt;211&gt; 22

&lt;212&gt; PRT

&lt;213&gt; Conus magus

&lt;220&gt;

&lt;221&gt; PEPTIDE

&lt;222&gt; (1)..(22)

<223> Xaa at residue 5 is Glu or gamma-carboxy Glu; Xaa at residue 8 and  
 10 is Pro or Hyp; Xaa at residue 12 is Tyr, 125I-Tyr, mono-iodo-  
 -Tyr, di-iodo-Tyr, O-sulpho-Tyr or O-phospho-Ty

&lt;400&gt; 142

Met Cys Cys Gly Xaa Ser Ala Xaa Cys Xaa Ser Xaa Phe Arg Asn Ser  
 1 5 10 15

Gln Ile Cys His Cys Cys  
 20

&lt;210&gt; 143

&lt;211&gt; 501

&lt;212&gt; DNA

&lt;213&gt; Conus magus

&lt;400&gt; 143

caagagggat cgatagcagt tcatgatgtc taaactggga gtcttggtga ccatctgtct 60  
 gcttctgttt ccccttactg ctcttccaat ggatggagat caacctgcag accaacctgc 120  
 agatcgtatg caggacgaca tttcatctga gcagtatccc ttgtttgata agagacaaaa 180  
 gtgttgcggc cccggcgggt catgccccgt atatttcaca gacaatttta tttgtggttg 240  
 ttgttaaagt acaactgtgc gatgaccaac ttcatatca cgactacgcc aagtgtctaa 300  
 tgaataaata aaatgattgc agtctcgctc agatttgctt ttgtatttg tctaaagatc 360  
 aatgacaaaa ccgtgtttt ggtgctggat tttcatatat ttctcgattc ctatccaaca 420  
 ctagatgatt taatcacgat agatctgatt tttttatcaa tgccttaatt ttttgctctg 480  
 tcatatcagt tttgtttata t 501

&lt;210&gt; 144

&lt;211&gt; 74

&lt;212&gt; PRT

43

&lt;213&gt; Conus magus

&lt;400&gt; 144

Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15  
 Pro Leu Thr Ala Leu Pro Met Asp Gly Asp Gln Pro Ala Asp Gln Pro  
 20 25 30  
 Ala Asp Arg Met Gln Asp Asp Ile Ser Ser Glu Gln Tyr Pro Leu Phe  
 35 40 45  
 Asp Lys Arg Gln Lys Cys Cys Gly Pro Gly Gly Ser Cys Pro Val Tyr  
 50 55 60  
 Phe Thr Asp Asn Phe Ile Cys Gly Cys Cys  
 65 70

&lt;210&gt; 145

&lt;211&gt; 23

&lt;212&gt; PRT

&lt;213&gt; Conus magus

&lt;220&gt;

&lt;221&gt; PEPTIDE

&lt;222&gt; (1)..(23)

<223> Xaa at residue 1 is Gln or pyro-Glu; Xaa at residue 6 and 11 is P  
 ro or Hyp; Xaa at residue 13 is Tyr, 125I-Tyr, mono-iodo-Tyr, di-  
 iodo-Tyr, O-sulpho-Tyr or O-phospho-Ty

&lt;400&gt; 145

Xaa Lys Cys Cys Gly Xaa Gly Gly Ser Cys Xaa Val Xaa Phe Thr Asp  
 1 5 10 15  
 Asn Phe Ile Cys Gly Cys Cys  
 20

&lt;210&gt; 146

&lt;211&gt; 454

&lt;212&gt; DNA

&lt;213&gt; Conus magus

&lt;400&gt; 146

caagagggat cgatagcagt tcatgatgtc taaactggga gtcttggtga ccatctgtct 60  
 gcttctgttt ccccttactg ctcttccaat ggatggagat caacctgcag accaacctgc 120  
 agatcgtatg caggacgaca ttcatctga gcagatccc ttgtttgata agagacaaaa 180  
 gtgttgcggc cccggcgggt catgccccgt atatttcaga gacaatttta tttgtggttg 240  
 ttgttaaatg acaacgtgtc gatgaccatc ttcattatca cgactacgcc aagtgtctaa 300  
 tgaataaata aaatgattgc agtctcgtc agatttgctt ttgtattttg gtctaaagat 360  
 caatgaccaa accgttggtt tgggtgtgat ttcatatat ttctcgattc ctatccaaca 420  
 ctagatgatt taatcacgat agatctgatt tttt 454

&lt;210&gt; 147

&lt;211&gt; 74

&lt;212&gt; PRT

&lt;213&gt; Conus magus

&lt;400&gt; 147

44

Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
1 5 10 15

Pro Leu Thr Ala Leu Pro Met Asp Gly Asp Gln Pro Ala Asp Gln Pro  
20 25 30

Ala Asp Arg Met Gln Asp Asp Ile Ser Ser Glu Gln Tyr Pro Leu Phe  
35 40 45

Asp Lys Arg Gln Lys Cys Cys Gly Pro Gly Gly Ser Cys Pro Val Tyr  
50 55 60

Phe Arg Asp Asn Phe Ile Cys Gly Cys Cys  
65 70

<210> 148

<211> 23

<212> PRT

<213> Conus magus

<220>

<221> PEPTIDE

<222> (1)..(23)

<223> Xaa at residue 1 is Gln or pyro-Glu; Xaa at residue 6 and 11 is P  
ro or Hyp; Xaa at residue 13 is Tyr, 125I-Tyr, mono-iodo-Tyr, di-  
iodo-Tyr, O-sulpho-Tyr or O-phospho-Ty

<400> 148

Xaa Lys Cys Cys Gly Xaa Gly Gly Ser Cys Xaa Val Xaa Phe Arg Asp  
1 5 10 15

Asn Phe Ile Cys Gly Cys Cys  
20

<210> 149

<211> 22

<212> PRT

<213> Conus magus

<220>

<221> PEPTIDE

<222> (1)..(22)

<223> Xaa at residue 1 is Gln or pyro-Glu; Xaa at residue 10 and 20 is  
Pro or Hyp; Xaa at residue 12 is Tyr, 125I-Tyr, mono-iodo-Tyr, di-  
iodo-Tyr, O-sulpho-Tyr or O-phospho-Ty

<400> 149

Xaa Lys Cys Cys Ser Gly Gly Ser Cys Xaa Leu Xaa Phe Arg Asp Arg  
1 5 10 15

Leu Ile Cys Xaa Cys Cys  
20

<210> 150

<211> 19

<212> PRT

<213> Conus marmoreus

<220>

<221> PEPTIDE

<222> (1)..(19)

<223> Xaa at residue 16 is Pro or Hyp

<400> 150

Ser Lys Gln Cys Cys His Leu Ala Ala Cys Arg. Phe Gly Cys Thr Xaa  
1 5 10 15

Cys Cys Asn

<210> 151  
 <211> 321  
 <212> DNA  
 <213> Conus marmoreus  
 <400> 151  
 caagaaggat cgatagcagt tcatgatgtc taaactggga gtcttggtga ccattctgtct 60  
 gcttctgttt cccgttactg ctcttcgat ggatggatgat caacctgcag accgacttgt 120  
 agagcgtatg caggacaaca ttcatctga gcagcatccc ttctttgaaa agagaagagg 180  
 aggctgttgc acacctccga ggaaatgcaa agaccgagcc tgcaaacctg cacgttgctg 240  
 cggcccagga taactgtgtg atgaccaact ttgttatcac ggctacgtca agtgtctagt 300  
 gaataagtaa aacgattgca g 321

<210> 152  
 <211> 76  
 <212> PRT  
 <213> Conus marmoreus  
 <400> 152  
 Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15  
 Pro Val Thr Ala Leu Pro Met Asp Gly Asp Gln Pro Ala Asp Arg Leu  
 20 25 30  
 Val Glu Arg Met Gln Asp Asn Ile Ser Ser Glu Gln His Pro Phe Phe  
 35 40 45  
 Glu Lys Arg Arg Gly Gly Cys Cys Thr Pro Pro Arg Lys Cys Lys Asp  
 50 55 60  
 Arg Ala Cys Lys Pro Ala Arg Cys Cys Gly Pro Gly  
 65 70 75

<210> 153  
 <211> 24  
 <212> PRT  
 <213> Conus marmoreus

<220>  
 <221> PEPTIDE  
 <222> (1)..(24)  
 <223> Xaa at residue 3, 8, 18 and 24 is Pro or Hyp

<400> 153  
 Arg Gly Gly Cys Cys Thr Xaa Xaa Arg Lys Cys Lys Asp Arg Ala Cys  
 1 5 10 15  
 Lys Xaa Ala Arg Cys Cys Gly Xaa  
 20

<210> 154  
 <211> 296  
 <212> DNA  
 <213> Conus marmoreus

<400> 154  
 gagctcggta ccccgacctc aagagggatc gatagcagtt catgatgtct aaactgggaa 60



46

tcttggtgac catctgtcta cttctatttc cccttactgc tgttccgctg gatggagatc 120  
 aacctgcaga ccgacctgca gagcgtatgc aggacgacat ttcacttgaa catcatccct 180  
 tttttgatcc cgtcaaacgg tgttgaggt tatcatgcgg cctgggatgc cacccttggt 240  
 gtggatgacc agctttgtta tcgcggcctc atcaagtgtc taatgaataa gtaaaa 296

<210> 155  
 <211> 68  
 <212> PRT  
 <213> Conus marmoreus

<400> 155  
 Met Met Ser Lys Leu Gly Ile Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15  
 Pro Leu Thr Ala Val Pro Leu Asp Gly Asp Gln Pro Ala Asp Arg Pro  
 20 25 30  
 Ala Glu Arg Met Gln Asp Asp Ile Ser Ser Glu His His Pro Phe Phe  
 35 40 45  
 Asp Pro Val Lys Arg Cys Cys Arg Leu Ser Cys Gly Leu Gly Cys His  
 50 55 60

Pro Cys Cys Gly  
 65

<210> 156  
 <211> 14  
 <212> PRT  
 <213> Conus marmoreus

<220>  
 <221> PEPTIDE  
 <222> (1)..(14)  
 <223> Xaa at residue 12 is Pro or Hyp

<400> 156  
 Cys Cys Arg Leu Ser Cys Gly Leu Gly Cys His Xaa Cys Cys  
 1 5 10

<210> 157  
 <211> 355  
 <212> DNA  
 <213> Conus marmoreus

<400> 157  
 ggcctacacc aagcttgcat gcctgcaggt cgactctaga ggatccccga tcgatagcag 60  
 ttcactgatgt ctgactggg agtcttggtg accatctgtc tacttctgtt tccccttact 120  
 gctgttccgc tggatggaga tcaacctgcg gaccgacctg cagagcgctt gcaggacgac 180  
 atttcatctg aacatcatcc ccattttgat tccggcagag agtggtgcgg ttcgttcgca 240  
 tgccgctttg gatgcgtgcc ttgttggtga tgaccagctt tgttatcacg gcctcatcga 300  
 gtgtctaatag aataagtaaa acgattgcag tagggcggta ccgagctcga attcc 355

<210> 158  
 <211> 69  
 <212> PRT

47

&lt;213&gt; Conus marmoreus

&lt;400&gt; 158

Met Met Ser Arg Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Phe  
 1 5 10 15

Pro Leu Thr Ala Val Pro Leu Asp Gly Asp Gln Pro Ala Asp Arg Pro  
 20 25 30

Ala Glu Arg Leu Gln Asp Asp Ile Ser Ser Glu His His Pro His Phe  
 35 40 45

Asp Ser Gly Arg Glu Cys Cys Gly Ser Phe Ala Cys Arg Phe Gly Cys  
 50 55 60

Val Pro Cys Cys Val  
 65

&lt;210&gt; 159

&lt;211&gt; 17

&lt;212&gt; PRT

&lt;213&gt; Conus marmoreus

&lt;220&gt;

&lt;221&gt; PEPTIDE

&lt;222&gt; (1)..(17)

<223> Xaa at residue 1 is Glu or gamma-carboxy Glu; Xaa at residue 14 i  
 s Pro or Hy

&lt;400&gt; 159

Xaa Cys Cys Gly Ser Phe Ala Cys Arg Phe Gly Cys Val Xaa Cys Cys  
 1 5 10 15

Val

&lt;210&gt; 160

&lt;211&gt; 295

&lt;212&gt; DNA

&lt;213&gt; Conus marmoreus

&lt;400&gt; 160

cgacctcaag agggatcgat agcagttcat gatgtctaaa ctgggagtct tgttgaccat 60

ctgtctactt ctatttcccc ttactgctgt tccgctggat ggagaccaac ctgcagaccg 120

acctgcagag cgtatgcagg acgacatttc atctgaacgt catccttttt ttgatcgcag 180

caaacagtgt tgccatctgc cggcatgccg cttcggatgt acgccttggt gttggtgatc 240

agctttgtta tcgcgtcttc atcaagtgtc taatgaataa gtaaaatgat tgcag 295

&lt;210&gt; 161

&lt;211&gt; 67

&lt;212&gt; PRT

&lt;213&gt; Conus marmoreus

&lt;400&gt; 161

Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Phe  
 1 5 10 15

Pro Leu Thr Ala Val Pro Leu Asp Gly Asp Gln Pro Ala Asp Arg Pro  
 20 25 30

Ala Glu Arg Met Gln Asp Asp Ile Ser Ser His Pro Phe Phe Asp Arg  
 35 40 45

Ser Lys Gln Cys Cys His Leu Pro Ala Cys Arg Phe Gly Cys Thr Pro  
 50 55 60

Cys Cys Trp  
 65

<210> 162  
 <211> 19  
 <212> PRT  
 <213> Conus marmoreus

<220>  
 <221> PEPTIDE  
 <222> (1)..(19)  
 <223> Xaa at residue 8 and 16 is Pro or Hyp; Xaa at residue 19 is Trp o  
 r bromo-Tr

<400> 162  
 Ser Lys Gln Cys Cys His Leu Xaa Ala Cys Arg Phe Gly Cys Thr Xaa  
 1 5 10 15

Cys Cys Xaa

<210> 163  
 <211> 235  
 <212> DNA  
 <213> Conus marmoreus

<400> 163  
 ggatccatga tgtctaaact gggagtcttg ttgaccatct gtctgcttct gtttcccctt 60  
 actgtctcttc cgctggatgg agatcaacct gcagaccaac gtgcagagcg tacgcaggcc 120  
 gagaagcatt ccttgccctga tccgagaatg ggctgttgcc cgtttccatg caaaaccagt 180  
 tgcactactt tgtgttgctg gtgatgataa cgtgttgatg accaactttc togag 235

<210> 164  
 <211> 67  
 <212> PRT  
 <213> Conus marmoreus

<400> 164  
 Gly Ser Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu  
 1 5 10 15

Leu Phe Pro Leu Thr Ala Leu Pro Leu Asp Gly Asp Gln Pro Ala Asp  
 20 25 30

Gln Arg Ala Glu Arg Thr Gln Ala Glu Lys His Ser Leu Pro Asp Pro  
 35 40 45

Arg Met Gly Cys Cys Pro Phe Pro Cys Lys Thr Ser Cys Thr Thr Leu  
 50 55 60

Cys Cys Gly  
 65

<210> 165  
 <211> 17  
 <212> PRT  
 <213> Conus marmoreus

<220>

49

&lt;221&gt; PEPTIDE

&lt;222&gt; (1)..(17)

&lt;223&gt; Xaa at residue 5 and 7 is Pro or Hyp

&lt;400&gt; 165

Met	Gly	Cys	Cys	Xaa	Phe	Xaa	Cys	Lys	Thr	Ser	Cys	Thr	Thr	Leu	Cys
1				5					10					15	

Cys

&lt;210&gt; 166

&lt;211&gt; 16

&lt;212&gt; PRT

&lt;213&gt; Conus marmoreus

&lt;220&gt;

&lt;221&gt; PEPTIDE

&lt;222&gt; (1)..(16)

&lt;223&gt; Xaa at residue 4 and 6 is Trp or bromo-Trp

&lt;400&gt; 166

Cys	Cys	His	Xaa	Asn	Xaa	Cys	Asp	His	Leu	Cys	Ser	Cys	Cys	Gly	Ser
1				5					10					15	

&lt;210&gt; 167

&lt;211&gt; 357

&lt;212&gt; DNA

&lt;213&gt; Conus marmoreus

&lt;400&gt; 167

gcc	aag	ctt	g	cat	gc	ct	g	ca	gg	at	g	act	t	ct	ag	agg	at	ccc	cac	ct	ca	aga	ggg	at	c	g	ata	60					
gc	agt	t	cat	g	at	gt	ct	aa	ac	tggg	ag	t	ct	t	gt	g	acc	at	c	t	g	t	ct	act	t	tc	tg	tt	gc	ct	120		
t	act	g	ct	gt	t	ccg	ct	gg	at	g	g	at	ca	acc	tgc	ag	acc	ga	c	ct	gc	aga	aac	g	t	at	gc	ag	ga	180			
cg	ac	att	t	ca	t	ct	ga	ac	gt	c	at	cc	at	gt	t	tg	at	gc	cg	t	c	ag	at	t	gt	t	g	cc	cc	gt	g	cc	240
gg	cat	g	cccc	t	tt	gg	at	g	ca	acc	ct	t	gt	gt	t	gg	at	ga	cca	g	ct	tt	gt	t	at	cg	gg	ac	ct	ca	300		
t	ca	ag	t	gt	ct	a	at	ga	ata	ag	t	aaaa	a	ac	ga	t	tc	g	ag	t	gg	t	acc	g	ag	ct	c	ga	att	cc	357		

&lt;210&gt; 168

&lt;211&gt; 67

&lt;212&gt; PRT

&lt;213&gt; Conus marmoreus

&lt;400&gt; 168

Met	Met	Ser	Lys	Leu	Gly	Val	Leu	Leu	Thr	Ile	Cys	Leu	Leu	Leu	Phe
1				5					10					15	

Ala	Leu	Thr	Ala	Val	Pro	Leu	Asp	Gly	Asp	Gln	Pro	Ala	Asp	Arg	Pro
			20					25					30		

Ala	Glu	Arg	Met	Gln	Asp	Asp	Ile	Ser	Ser	His	Pro	Met	Phe	Asp	Ala
			35				40					45			

Val	Arg	Asp	Cys	Cys	Pro	Leu	Pro	Ala	Cys	Pro	Phe	Gly	Cys	Asn	Pro
			50			55					60				

Cys	Cys	Gly
65		

&lt;210&gt; 169

&lt;211&gt; 16

50

<212> PRT  
<213> Conus marmoreus

<220>  
<221> PEPTIDE  
<222> (1)..(16)  
<223> Xaa at residue 4, 6, 9 and 14 is Pro or Hyp

<400> 169  
Asp Cys Cys Xaa Leu Xaa Ala Cys Xaa Phe Gly Cys Asn Xaa Cys Cys  
1 5 10 15

<210> 170  
<211> 16  
<212> PRT  
<213> Conus marmoreus

<220>  
<221> PEPTIDE  
<222> (1)..(16)  
<223> Xaa at residue 4 and 13 is Pro or Hyp

<400> 170  
Cys Cys Ala Xaa Ser Ala Cys Arg Leu Gly Cys Arg Xaa Cys Cys Arg  
1 5 10 15

<210> 171  
<211> 16  
<212> PRT  
<213> Conus marmoreus

<220>  
<221> PEPTIDE  
<222> (1)..(16)  
<223> Xaa at residue 4 and 13 is Pro or Hyp

<400> 171  
Cys Cys Ala Xaa Ser Ala Cys Arg Leu Gly Cys Arg Xaa Cys Cys Arg  
1 5 10 15

<210> 172  
<211> 16  
<212> PRT  
<213> Conus marmoreus

<220>  
<221> PEPTIDE  
<222> (1)..(16)  
<223> Xaa at residue 4 and 13 is Pro or Hyp

<400> 172  
Cys Cys Ala Xaa Ser Ala Cys Arg Leu Gly Cys Arg Xaa Cys Cys Arg  
1 5 10 15

<210> 173  
<211> 17  
<212> PRT  
<213> Conus marmoreus

<220>  
<221> PEPTIDE  
<222> (1)..(17)  
<223> Xaa at residue 14 is Pro or Hyp

<400> 173

51.

Gly Cys Cys Gly Ser Phe Ala Cys Arg Phe Gly Cys Val Xaa Cys Cys  
 1 5 10 15

Val

<210> 174  
 <211> 244  
 <212> DNA  
 <213> Conus nobilis

<400> 174  
 ggatccatga tgtctaaact gggagtcttg ttgaccatct gtctacttct gtttcccctt 60  
 actgctcttc cgctggatga agatcaaccg gtacaccgac ctgcagagcg tatgcaggac 120  
 atttcatctg atcaacatct cttctttgat ctcatcaaac ggtgctgcga gttgccatgc 180  
 gggccaggct tttgcgtccc ttgttgctga catcaataac gtgttgatga ccaactttct 240  
 cgag 244

<210> 175  
 <211> 69  
 <212> PRT  
 <213> Conus nobilis

<400> 175  
 Gly Ser Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu  
 1 5 10 15  
 Leu Phe Pro Leu Thr Ala Leu Pro Leu Asp Glu Asp Gln Pro Val His  
 20 25 30  
 Arg Pro Ala Glu Arg Met Gln Asp Ile Ser Ser Asp Gln His Leu Phe  
 35 40 45  
 Phe Asp Leu Ile Lys Arg Cys Cys Glu Leu Pro Cys Gly Pro Gly Phe  
 50 55 60

Cys Val Pro Cys Cys  
 65

<210> 176  
 <211> 15  
 <212> PRT  
 <213> Conus nobilis

<220>  
 <221> PEPTIDE  
 <222> (1)..(15)  
 <223> Xaa at residue 3 is Glu or gamma-carboxy Glu; Xaa at residue 5, 8  
 adn 13 is Pro or Hy

<400> 176  
 Cys Cys Xaa Leu Xaa Cys Gly Xaa Gly Phe Cys Val Xaa Cys Cys  
 1 5 10 15

<210> 177  
 <211> 262  
 <212> DNA  
 <213> Conus nobilis

<400> 177  
 ggatccatga tgtctaaact gggagtcttg ttgaccatct gtctacttct gtttcccctt 60

52

actgcttttc cgatggatgg agatcaacct gcagaccaac ctgcagatcg tatgcaggac 120  
 gacatttcat ctgagcagta tcccttggtt gataagagac aaaagtgttg cactgggaag 180  
 aaggggtcat gctccggcaa agcatgcaaa aatctcaa atgtgtcttg acgataacgt 240  
 gttgatgacc aactttctcg ag 262

<210> 178  
 <211> 78  
 <212> PRT  
 <213> Conus nobilis

<400> 178  
 Gly Ser Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu  
 1 5 10 15  
 Leu Phe Pro Leu Thr Ala Phe Pro Met Asp Gly Asp Gln Pro Ala Asp  
 20 25 30  
 Gln Pro Ala Asp Arg Met Gln Asp Asp Ile Ser Ser Glu Gln Tyr Pro  
 35 40 45  
 Leu Phe Asp Lys Arg Gln Lys Cys Cys Thr Gly Lys Lys Gly Ser Cys  
 50 55 60  
 Ser Gly Lys Ala Cys Lys Asn Leu Lys Cys Cys Ser Gly Arg  
 65 70 75

<210> 179  
 <211> 23  
 <212> PRT  
 <213> Conus nobilis

<220>  
 <221> PEPTIDE  
 <222> (1)..(23)  
 <223> Xaa at residue 1 is Gln or pyro-Glu

<400> 179  
 Xaa Lys Cys Cys Thr Gly Lys Lys Gly Ser Cys Ser Gly Lys Ala Cys  
 1 5 10 15  
 Lys Asn Leu Lys Cys Cys Ser  
 20

<210> 180  
 <211> 238  
 <212> DNA  
 <213> Conus pulicarius

<400> 180  
 ggatccatga tgtctaaact gggagttttg ttgaccatct gtctgcttct gtttcccctt 60  
 actgctgttc cgctggatgg agatcaacct gcagaccgac ctgcagagcg tatgcaggac 120  
 attgcaactg aacagcatcc cttctttgat cccgtcaaac ggtgttgcaa cagctgttac 180  
 atgggatgca tcccttggtg cttctagtaa taacgtgttg atgaccaact ttctcgag 238

<210> 181  
 <211> 68  
 <212> PRT  
 <213> Conus pulicarius

53

&lt;400&gt; 181

Gly Ser Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu  
 1 5 10 15

Leu Phe Pro Leu Thr Ala Val Pro Leu Asp Gly Asp Gln Pro Ala Asp  
 20 25 30

Arg Pro Ala Glu Arg Met Gln Asp Ile Ala Thr Glu Gln His Pro Phe  
 35 40 45

Phe Asp Pro Val Lys Arg Cys Cys Asn Ser Cys Tyr Met Gly Cys Ile  
 50 55 60

Pro Cys Cys Phe  
 65

&lt;210&gt; 182

&lt;211&gt; 14

&lt;212&gt; PRT

&lt;213&gt; Conus pulicarius

&lt;220&gt;

&lt;221&gt; PEPTIDE

&lt;222&gt; (1)..(14)

<223> Xaa at residue 11 is Pro or Hyp; Xaa at residue 5 is Tyr, 125I-Ty  
 r, mono-iodo-Tyr, di-iodo-Tyr, O-sulpho-Tyr or O-phospho-Ty

&lt;400&gt; 182

Cys Cys Asn Ser Cys Xaa Met Gly Cys Ile Xaa Cys Cys Phe  
 1 5 10

&lt;210&gt; 183

&lt;211&gt; 238

&lt;212&gt; DNA

&lt;213&gt; Conus quercinus

&lt;400&gt; 183

ggatccatga tgtctaaact gggagtcttg ttgaccatct gtctgcttct gtttccctt 60

acagctcttc agctggatgg agatcaacct gcagaccgac ctgcagagcg tacgcaggac 120

attgcatctg aacagtatcg aaagtttgat cagagacaga ggtgttgcca gtggccatgc 180

cccggtagtt gcagatgctg ccgtactggt taacgtgttg atgaccaact ttctcgag 238

&lt;210&gt; 184

&lt;211&gt; 70

&lt;212&gt; PRT

&lt;213&gt; Conus quercinus

&lt;400&gt; 184

Gly Ser Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu  
 1 5 10 15

Leu Phe Pro Leu Thr Ala Leu Gln Leu Asp Gly Asp Gln Pro Ala Asp  
 20 25 30

Arg Pro Ala Glu Arg Thr Gln Asp Ile Ala Ser Glu Gln Tyr Arg Lys  
 35 40 45

Phe Asp Gln Arg Gln Arg Cys Cys Gln Trp Pro Cys Pro Gly Ser Cys  
 50 55 60

Arg Cys Cys Arg Thr Gly  
 65 70



<210> 185  
 <211> 17  
 <212> PRT  
 <213> Conus quercinus

<220>  
 <221> PEPTIDE  
 <222> (1)..(17)  
 <223> Xaa at residue 1 is Gln or pyro-Glu; Xaa at residue 7 and 9 is Pro or Hyp; Xaa at residue 6 is Trp or bromo-Trp

<400> 185  
 Xaa Arg Cys Cys Gln Xaa Xaa Cys Xaa Gly Ser Cys Arg Cys Cys Arg  
 1 5 10 15

Thr

<210> 186  
 <211> 15  
 <212> PRT  
 <213> Conus quercinus

<220>  
 <221> PEPTIDE  
 <222> (1)..(15)  
 <223> Xaa at residue 11 and 14 is Pro or Hyp

<400> 186  
 Cys Cys Ser Gln Asp Cys Leu Val Cys Ile Xaa Cys Cys Xaa Asn  
 1 5 10 15

<210> 187  
 <211> 15  
 <212> PRT  
 <213> Conus quercinus

<220>  
 <221> PEPTIDE  
 <222> (1)..(15)  
 <223> Xaa at residue 11 14 is Pro or Hyp; Xaa at residue 7 is Trp or bromo-Trp

<400> 187  
 Cys Cys Ser Arg His Cys Xaa Val Cys Ile Xaa Cys Cys Xaa Asn  
 1 5 10 15

<210> 188  
 <211> 323  
 <212> DNA  
 <213> Conus radiatus

<400> 188  
 tcaagaagga tcgatagcag ttcgatgatgt ctaaactggg agtcttgttg accatctgtc 60  
 tgcttctgtt tccccttact gctcttcga tggatggaga tcaacctgta gaccgacttg 120  
 cagagcgtat gcaggacaac atttcactctg agcagcatac cttctttgaa aagagactac 180  
 catcgtgttg ctcccttaac ttgcggcttt gccagtagc agcatgcaaa cgtaaccctt 240  
 gttgcacagg ataactgtgtt gatgaccaac tttgttatca cggtacgtc aagtgtctag 300  
 tgaataagta aaacgattgc agt 323

55

<210> 189  
 <211> 76  
 <212> PRT  
 <213> Conus radiatus

<400> 189  
 Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15  
 Pro Leu Thr Ala Leu Pro Met Asp Gly Asp Gln Pro Val Asp Arg Leu  
 20 25 30  
 Ala Glu Arg Met Gln Asp Asn Ile Ser Ser Glu Gln His Thr Phe Phe  
 35 40 45  
 Glu Lys Arg Leu Pro Ser Cys Cys Ser Leu Asn Leu Arg Leu Cys Pro  
 50 55 60  
 Val Pro Ala Cys Lys Arg Asn Pro Cys Cys Thr Gly  
 65 70 75

<210> 190  
 <211> 24  
 <212> PRT  
 <213> Conus radiatus

<220>  
 <221> PEPTIDE  
 <222> (1)..(24)  
 <223> Xaa at residue 2, 13, 15 and 21 is Pro or Hyp

<400> 190  
 Leu Xaa Ser Cys Cys Ser Leu Asn Leu Arg Leu Cys Xaa Val Xaa Ala  
 1 5 10 15

Cys Lys Arg Asn Xaa Cys Cys Thr  
 20

<210> 191  
 <211> 336  
 <212> DNA  
 <213> Conus radiatus

<400> 191  
 aggtcgactc tagaggatcc ccaaggatcg atagcagttc atgatgtcta aactgggagt 60  
 cttgttgacc atctgtotgc ttctgtttcc ccttactgct cttccgatgg atggagatca 120  
 acctgcagac cgacttgacag agcgtatgca ggacgacatt tcatctgagc agcatccott 180  
 ctttaaaaag agacaacaaa gatgttgacac cgttaagagg atttgtccag taccagcatg 240  
 cagaagtaaa cttgttgca aatcataacg tattgatgac caactttgtt atcacggcta 300  
 cgtcaagtgt ctagtgaata agtaaaatga ttgcag 336

<210> 192  
 <211> 75  
 <212> PRT  
 <213> Conus radiatus

<400> 192  
 Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15

56

Pro Leu Thr Ala Leu Pro Met Asp Gly Asp Gln Pro Ala Asp Arg Leu  
20 25 30

Ala Glu Arg Met Gln Asp Asp Ile Ser Ser Glu Gln His Pro Phe Phe  
35 40 45

Lys Lys Arg Gln Gln Arg Cys Cys Thr Val Lys Arg Ile Cys Pro Val  
50 55 60

Pro Ala Cys Arg Ser Lys Pro Cys Cys Lys Ser  
65 70 75

<210> 193

<211> 24

<212> PRT

<213> Conus radiatus

<220>

<221> PEPTIDE

<222> (1)..(24)

<223> Xaa at residue 1 is Gln or pyro-Glu; Xaa at residue 12, 14 and 20  
is Pro or Hy

<400> 193

Xaa Gln Arg Cys Cys Thr Val Lys Arg Ile Cys Xaa Val Xaa Ala Cys  
1 5 10 15

Arg Ser Lys Xaa Cys Cys Lys Ser  
20

<210> 194

<211> 326

<212> DNA

<213> Conus radiatus

<400> 194

acctcaagaa ggatcgatag cagttcatga tgtctaaact gggagtcttg ttgaccatct 60  
gtctgcttct gtttcccggtt actgctcttc cgatggatgg tgatcaacct gcagaccgac 120  
ttgtagagcg tatgcaggac aacatttcat ctgagcagca tcccttcttt gaaaagagaa 180  
gaggaggctg ttgcacacct ccgaggaaat gcaaagaccg agcctgcaaa cctgcacgtt 240  
gctgcggccc aggataacgt gttgatgacc aactttgtta tcacggctac gtcaagtgtc 300  
tagtgaataa gtaaaacgat tgcagt 326

<210> 195

<211> 76

<212> PRT

<213> Conus radiatus

<400> 195

Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
1 5 10 15

Pro Val Thr Ala Leu Pro Met Asp Gly Asp Gln Pro Ala Asp Arg Leu  
20 25 30

Val Glu Arg Met Gln Asp Asn Ile Ser Ser Glu Gln His Pro Phe Phe  
35 40 45

Glu Lys Arg Arg Gly Gly Cys Cys Thr Pro Pro Arg Lys Cys Lys Asp  
50 55 60

Arg Ala Cys Lys Pro Ala Arg Cys Cys Gly Pro Gly  
65 70 75

<210> 196  
<211> 24  
<212> PRT  
<213> Conus radiatus

<220>  
<221> PEPTIDE  
<222> (1)..(24)  
<223> Xaa at residue 7, 8, 18 and 24 is Pro or Hyp

<400> 196  
Arg Gly Gly Cys Cys Thr Xaa Xaa Arg Lys Cys Lys Asp Arg Ala Cys  
1 5 10 15

Lys Xaa Ala Arg Cys Cys Gly Xaa  
20

<210> 197  
<211> 238  
<212> DNA  
<213> Conus rattus

<400> 197  
ggatccatga tgtctaaact gggagtcttg gtgaccatct gctgcttct gttccctctt 60  
gctgcttttc cactggatgg agatcaacct gcagaccacc ctgcaaagcg tacgcaagat 120  
gacagttcag ctgccctgat caatgcctgg ctgatgaat cccagacttg ctgcagtaac 180  
tgcggtgaag attgtgatgg ttgttgccag taacgtgttg atgaccaact ttctcgag 238

<210> 198  
<211> 70  
<212> PRT  
<213> Conus rattus

<400> 198  
Gly Ser Met Met Ser Lys Leu Gly Val Leu Val Thr Ile Cys Leu Leu  
1 5 10 15

Leu Phe Pro Leu Ala Ala Phe Pro Leu Asp Gly Asp Gln Pro Ala Asp  
20 25 30

His Pro Ala Lys Arg Thr Gln Asp Asp Ser Ser Ala Ala Leu Ile Asn  
35 40 45

Ala Trp Leu Asp Glu Ser Gln Thr Cys Cys Ser Asn Cys Gly Glu Asp  
50 55 60

Cys Asp Gly Cys Cys Gln  
65 70

<210> 199  
<211> 16  
<212> PRT  
<213> Conus rattus

<220>  
<221> PEPTIDE  
<222> (1)..(16)  
<223> Xaa at residue 1 is Gln or pyro-Glu; Xaa at residue 9 is Glu or gamma-carboxy Gl

&lt;400&gt; 199

Xaa Thr Cys Cys Ser Asn Cys Gly Xaa Asp Cys Asp Gly Cys Cys Gln  
 1 5 10 15

&lt;210&gt; 200

&lt;211&gt; 327

&lt;212&gt; DNA

&lt;213&gt; Conus stercusmuscarum

&lt;400&gt; 200

gacctcaaga gggatcgata gcagttcgtg atgtotaaac tgggagtcctt gttgaccatc 60  
 tgtctgcttc tgtttctct tactgctctt ccgatggatg gagatcaacc tgcagaccaa 120  
 cctgcagatc gtatgcagga cgacatttca tctgagcagt atcccttggt tgataagaga 180  
 caaaagtgtt gcactgggaa gaaggggtca tgctccggca aagcatgcaa aaatctcaaa 240  
 tgttgctctg gacgataacg tgttgatgac caactttggt atcacggcta cgtcaagtgt 300  
 ctaatgaata agtaaaacga ttgcagt 327

&lt;210&gt; 201

&lt;211&gt; 75

&lt;212&gt; PRT

&lt;213&gt; Conus stercusmuscarum

&lt;400&gt; 201

Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe Pro  
 1 5 10 15  
 Leu Thr Ala Leu Pro Met Asp Gly Asp Gln Pro Ala Asp Gln Pro Ala  
 20 25 30  
 Asp Arg Met Gln Asp Asp Ile Ser Ser Glu Gln Tyr Pro Leu Phe Asp  
 35 40 45  
 Lys Arg Gln Lys Cys Cys Thr Gly Lys Lys Gly Ser Cys Ser Gly Lys  
 50 55 60  
 Ala Cys Lys Asn Leu Lys Cys Cys Ser Gly Arg  
 65 70 75

&lt;210&gt; 202

&lt;211&gt; 23

&lt;212&gt; PRT

&lt;213&gt; Conus stercusmuscarum

&lt;220&gt;

&lt;221&gt; PEPTIDE

&lt;222&gt; (1)..(23)

&lt;223&gt; Xaa at residue 1 is Gln or pyro-Glu

&lt;400&gt; 202

Xaa Lys Cys Cys Thr Gly Lys Lys Gly Ser Cys Ser Gly Lys Ala Cys  
 1 5 10 15

Lys Asn Leu Lys Cys Cys Ser  
 20

&lt;210&gt; 203

&lt;211&gt; 316

&lt;212&gt; DNA

&lt;213&gt; Conus stercusmuscarum

<400> 203  
 gatcgatagc agttcgtgat gtctaaactg ggagtcttgt tgaccatctg tctgcttctg 60  
 tttccctta ctgctcttcc gatggatgga gatcaacctg cagaccaacc tgcagatcgt 120  
 atgcagaacg acatttcacg tgagcagtat cccttgtttg ataagagaca aaagtgttgc 180  
 ggccccggcg cgtcatgccc cagatatttc aaagacaatt ttatttggg ttgttgtaa 240  
 atgacaacgt gtgatgacc aacttcgtta tcacgacttc gccaaagtgc taatgaataa 300  
 gtaaaacgat tgcagt 316  
 <210> 204  
 <211> 73  
 <212> PRT  
 <213> *Conus stercusmuscarum*  
  
 <400> 204  
 Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe Pro  
 1 5 10 15  
 Leu Thr Ala Leu Pro Met Asp Gly Asp Gln Pro Ala Asp Gln Pro Ala  
 20 25 30  
 Asp Arg Met Gln Asn Asp Ile Ser Ser Glu Gln Tyr Pro Leu Phe Asp  
 35 40 45  
 Lys Arg Gln Lys Cys Cys Gly Pro Gly Ala Ser Cys Pro Arg Tyr Phe  
 50 55 60  
 Lys Asp Asn Phe Ile Cys Gly Cys Cys  
 65 70  
 <210> 205  
 <211> 23  
 <212> PRT  
 <213> *Conus stercusmuscarum*  
  
 <220>  
 <221> PEPTIDE  
 <222> (1)..(23)  
 <223> Xaa at residue 1 is Gln or pyro-Glu; Xaa at residue 6 and 11 is P  
 ro or Hyp; Xaa at residue 13 is Tyr, 125I-Tyr, mono-iodo-Tyr, di-  
 iodo-Tyr, O-sulpho-Tyr or O-phospho-Ty  
  
 <400> 205  
 Xaa Lys Cys Cys Gly Xaa Gly Ala Ser Cys Xaa Arg Xaa Phe Lys Asp  
 1 5 10 15  
 Asn Phe Ile Cys Gly Cys Cys  
 20  
  
 <210> 206  
 <211> 331  
 <212> DNA  
 <213> *Conus striatus*  
  
 <400> 206  
 cgacctttca agaggggatcg atagcagttc gcgatgtcta aactgggggt attgttgacc 60  
 atctgtctgc ttctgtttcc ccttactgct cttccgatgg atgaagatca acctgcagac 120  
 caacttgaag atcgtatgca ggacgacatt tcatctgagc agtatccctc gtttgtagg 180  
 agacaaaagt gttgcggcga aggctcgtca tgccccaaat atttcaaaaa caattttatt 240

60

tgtggttggt gttaaatgac aacgtgtcga tgaccaactt cgttatcacg actacgcaa 300

gtgtcttgtc taatgataat aaaatgattc c 331

<210> 207

<211> 73

<212> PRT

<213> Conus striatus

<400> 207

Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe Pro  
1 5 10 15

Leu Thr Ala Leu Pro Met Asp Glu Asp Gln Pro Ala Asp Gln Leu Glu  
20 25 30

Asp Arg Met Gln Asp Asp Ile Ser Ser Glu Gln Tyr Pro Ser Phe Val  
35 40 45

Arg Arg Gln Lys Cys Cys Gly Glu Gly Ser Ser Cys Pro Lys Tyr Phe  
50 55 60

Lys Asn Asn Phe Ile Cys Gly Cys Cys  
65 70

<210> 208

<211> 23

<212> PRT

<213> Conus striatus

<220>

<221> PEPTIDE

<222> (1)..(23)

<223> Xaa at residue 1 is Gln or pyro-Glu; Xaa at residue 6 is Glu or gamma-carboxy Glu; Xaa at residue 11 is Pro or Hyp; Xaa at residue 13 is Tyr, 125I-Tyr, mono-iodo-Tyr, di-iodo-Tyr, O-sulpho-Tyr or O-phospho-Ty

<400> 208

Xaa Lys Cys Cys Gly Xaa Gly Ser Ser Cys Xaa Lys Xaa Phe Lys Asn  
1 5 10 15

Asn Phe Ile Cys Gly Cys Cys  
20

<210> 209

<211> 256

<212> DNA

<213> Conus striatus

<400> 209

ggatccatga tgtctaaact gggagtcttg ttgaccgtct gtctgcttct gtttcccctt 60

actgctcttc cgctggatgg agatcaacct gcagaccgac ctgcagagcg tatgcaggac 120

gacatttcat ctgacgagca tcccttggtt gataagagac aaaactgttg caatggggga 180

tgctccagca aatggtgcag agatcacgca cgttgttgcg gtcgatgata acgtgttgat 240

gaccaacttt ctcgag 256

<210> 210

<211> 75

<212> PRT

<213> *Conus striatus*

<400> 210

Gly Ser Met Met Ser Lys Leu Gly Val Leu Leu Thr Val Cys Leu Leu  
1 5 10 15

Leu Phe Pro Leu Thr Ala Leu Pro Leu Asp Gly Asp Gln Pro Ala Asp  
20 25 30

Arg Pro Ala Glu Arg Met Gln Asp Asp Ile Ser Ser Asp Glu His Pro  
35 40 45

Leu Phe Asp Lys Arg Gln Asn Cys Cys Asn Gly Gly Cys Ser Ser Lys  
50 55 60

Trp Cys Arg Asp His Ala Arg Cys Cys Gly Arg  
65 70 75

<210> 211

<211> 20

<212> PRT

<213> *Conus striatus*

<220>

<221> PEPTIDE

<222> (1)..(20)

<223> Xaa at residue 1 is Gln or pyro-Glu; Xaa at residue 12 is Trp or  
bromo-Tr

<400> 211

Xaa Asn Cys Cys Asn Gly Gly Cys Ser Ser Lys Xaa Cys Arg Asp His  
1 5 10 15

Ala Arg Cys Cys  
20

<210> 212

<211> 235

<212> DNA

<213> *Conus tessulatus*

<400> 212

ggatccatga tgtctaaact gggagtcttg ttgaccatgt gtctgcttct gtttccctt 60

actgctgttc cgctggatgg agatcaacct gcagaccgac ctgcagagcg taggcaggac 120

attgcaactg acgatcatcc ttgtttgat cccgtcaaac ggtgctgcc aaatgctat 180

atgggatgca tcccttggtg catttagtaa cgtgttgatg accaactttc togag 235

<210> 213

<211> 68

<212> PRT

<213> *Conus tessulatus*

<400> 213

Gly Ser Met Met Ser Lys Leu Gly Val Leu Leu Thr Met Cys Leu Leu  
1 5 10 15

Leu Phe Pro Leu Thr Ala Val Pro Leu Asp Gly Asp Gln Pro Ala Asp  
20 25 30

Arg Pro Ala Glu Arg Arg Gln Asp Ile Ala Thr Asp Asp His Pro Leu  
35 40 45



62

Phe Asp Pro Val Lys Arg Cys Cys His Lys Cys Tyr Met Gly Cys Ile  
 50 55 60

Pro Cys Cys Ile  
 65

<210> 214  
 <211> 14  
 <212> PRT  
 <213> Conus tessulatus

<220>  
 <221> PEPTIDE  
 <222> (1)..(14)  
 <223> Xaa at residue 11 is Pro or Hyp; Xaa at residue 6 is Tyr, 125I-Tyr, mono-iodo-Tyr, di-iodo-Tyr, O-sulpho-Tyr or O-phospho-Ty

<400> 214  
 Cys Cys His Lys Cys Xaa Met Gly Cys Ile Xaa Cys Cys Ile  
 1 5 10

<210> 215  
 <211> 238  
 <212> DNA  
 <213> Conus tessulatus

<400> 215  
 ggatccatga tgtctaaact gggagtcttg ttgaccatct gtgtgcttct gtttcccctt 60  
 actgctgttc cgctggatgg agatcaacct gcagaccaac ctgcagagcg tacgcagaac 120  
 gaggcagcatc ccttgatga tcagaaaaga aagtgttgcc ggccgccatg cgccatgagc 180  
 tgcggcatgg ctagggtgtg ctattaatga taacgtgttg atgaccaact ttctcgag 238

<210> 216  
 <211> 68  
 <212> PRT  
 <213> Conus tessulatus

<400> 216  
 Gly Ser Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Val Leu  
 1 5 10 15

Leu Phe Pro Leu Thr Ala Val Pro Leu Asp Gly Asp Gln Pro Ala Asp  
 20 25 30

Gln Pro Ala Glu Arg Thr Gln Asn Glu Gln His Pro Leu Tyr Asp Gln  
 35 40 45

Lys Arg Lys Cys Cys Arg Pro Pro Cys Ala Met Ser Cys Gly Met Ala  
 50 55 60

Arg Cys Cys Tyr  
 65

<210> 217  
 <211> 18  
 <212> PRT  
 <213> Conus tessulatus

<220>  
 <221> PEPTIDE  
 <222> (1)..(18)  
 <223> Xaa at residue 5 and 6 is Pro or Hyp; Xaa at residue 18 is Tyr, 1

25I-Tyr, mono-iodo-Tyr, di-iodo-Tyr, O-sulpho-Tyr or O-phospho-Ty

<400> 217

Lys Cys Cys Arg Xaa Xaa Cys Ala Met Ser Cys Gly Met Ala Arg Cys  
1 5 10 15

Cys Xaa

<210> 218

<211> 564

<212> DNA

<213> Conus textile

<400> 218

gagtcaaccc actgtcacgc caagagcggg cgccacagct aaggcaagaa ggatcgatag 60  
cagttcatga tgtctaaact gggagccttg ttgaccatct gtctacttct gttttccctt 120  
actgtgttgc cgctggatgg agatcaacat gcagaccaac ctgcacagcg tctgcaggac 180  
cgcattccaa ctgaagatca tcccttattt gatcccaaca aacgggtgttg cccgccggtg 240  
gcatgcaaca tgggatgcaa gccttgttgt ggatgaccag ctttgttata gcggtctcat 300  
gaagtgtcta atgaataagt aaaacgattg cagtttcggt cagatttgct gttgtatttt 360  
gggtctaaaga ttaatgacca aactgttctt ttgatccgga ttttcacgta tttctcgatt 420  
cctattcaac actagataag ttaatcacga cagatctgat tttccatcaa tgccttgctt 480  
tttggctctgt catataaatc ttgtttatat ttaatttctc gtcactttca acacgcacac 540  
acacacacac acacacgcgc gcgc 564

<210> 219

<211> 69

<212> PRT

<213> Conus textile

<400> 219

Met Met Ser Lys Leu Gly Ala Leu Leu Thr Ile Cys Leu Leu Leu Phe  
1 5 10 15

Ser Leu Thr Ala Val Pro Leu Asp Gly Asp Gln His Ala Asp Gln Pro  
20 25 30

Ala Gln Arg Leu Gln Asp Arg Ile Pro Thr Glu Asp His Pro Leu Phe  
35 40 45

Asp Pro Asn Lys Arg Cys Cys Pro Pro Val Ala Cys Asn Met Gly Cys  
50 55 60

Lys Pro Cys Cys Gly  
65

<210> 220

<211> 16

<212> PRT

<213> Conus textile

<220>

<221> PEPTIDE

<222> (1)..(16)

<223> Xaa at residue 3, 4 and 13 is Pro or Hyp

64

&lt;400&gt; 220

Cys Cys Xaa Xaa Val Ala Cys Asn Met Gly Cys Lys Xaa Cys Cys Gly  
 1 5 10 15

&lt;210&gt; 221

&lt;211&gt; 780

&lt;212&gt; DNA

&lt;213&gt; Conus textile

&lt;400&gt; 221

ggatccagac gacaagaag agtcaacca ctgccacgtc aagagcagag cccacagcta 60  
 agacaagaag gatcgatagc agttcatgat gtttaaactg ggagtcttgt tgaccatctg 120  
 tctccttctg ttttccctta atgctgttcc gttggatgga gatcaacctg cagaccaacc 180  
 tgcagagcgt ctgctggacg acatttcatt tgaaaataat cccttttatg atcccgccaa 240  
 acggtgttgc aggacttgct tcggttgcac accttgttgt ggatgaccag cctcatcaag 300  
 tgtctaacga ataagtaaag cgattgcagt ctggttcaga tttacttttg tattctgggc 360  
 taaagattaa tgaccaaact cttcttttga tccggatgta catatatttc tcgattccta 420  
 tccaacgcta gataagctaa tcacgacaga tctgattttc tgtcaatgcc ttgctttttg 480  
 gtctctcata tcaactottgt ttatatttaa tttctcgtca ctatatatat atatacacac 540  
 acacacacac ggaattccga ttgtccagta ccgttcttgg gatcgaggta ttgctgcat 600  
 ggcttattct gtactctttt cttctgcgtg tgatagtgat gtcttctact cccatctgtg 660  
 ctacccttgg cttgatcttt gataggcgtg tgccttcac tgggtataaa cccctctgat 720  
 cctactctct ggacgcctcg ggggcccaac ctccaaataa agcgacatcc aatgaaaaaa 780

&lt;210&gt; 222

&lt;211&gt; 66

&lt;212&gt; PRT

&lt;213&gt; Conus textile

&lt;400&gt; 222

Met Met Phe Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15

Ser Leu Asn Ala Val Pro Leu Asp Gly Asp Gln Pro Ala Asp Gln Pro  
 20 25 30

Ala Glu Arg Leu Leu Asp Asp Ile Ser Phe Glu Asn Asn Pro Phe Tyr  
 35 40 45

Asp Pro Ala Lys Arg Cys Cys Arg Thr Cys Phe Gly Cys Thr Pro Cys  
 50 55 60

Cys Gly  
 65

&lt;210&gt; 223

&lt;211&gt; 12

&lt;212&gt; PRT

&lt;213&gt; Conus textile

&lt;220&gt;

&lt;221&gt; PEPTIDE

65

<222> (1)..(12)  
 <223> Xaa at residue 10 is Pro or Hyp  
  
 <400> 223  
 Cys Cys Arg Thr Cys Phe Gly Cys Thr Xaa Cys Cys  
 1 5 10  
  
 <210> 224  
 <211> 456  
 <212> DNA  
 <213> Conus textile  
  
 <400> 224  
 ggaacagtca accccacagc cagccaaga gcagacagcc acagctacgt gaagaagggt 60  
 ggagagaggt tcatgatgtt gaaaatggga gtggtgctat tcatctttct ggtactgttt 120  
 cccctggcaa cgctccagct ggatgcagat caacctgtag aacgatatgc ggagaacaaa 180  
 cagctcctca acccagatga aaggagggaa atcctattgc ctgctctgag gaagttctgc 240  
 tgtgattcga attggtgccca ctttcggat tgtgagtgcct gctacgggta gcgccgaaca 300  
 tccatggcac tgtgctgggc ggtttcatcc caacaacgac agcgtttgtt gatttcattgt 360  
 atcattgcgc ccacgtctct tgtctaagaa tgacgaacat gattgcactc tgggtcagat 420  
 ttcgtgttct tttctgacaa taaatgacaa acctcc 456  
  
 <210> 225  
 <211> 70  
 <212> PRT  
 <213> Conus textile  
  
 <400> 225  
 Met Met Leu Lys Met Gly Val Val Leu Phe Ile Phe Leu Val Leu Phe  
 1 5 10 15  
 Pro Leu Ala Thr Leu Gln Leu Asp Ala Asp Gln Pro Val Glu Arg Tyr  
 20 25 30  
 Ala Glu Asn Lys Gln Leu Leu Asn Pro Asp Glu Arg Arg Glu Ile Leu  
 35 40 45  
 Leu Pro Ala Leu Arg Lys Phe Cys Cys Asp Ser Asn Trp Cys His Asp  
 50 55 60  
 Cys Glu Cys Cys Tyr Gly  
 65 70  
 <210> 226  
 <211> 17  
 <212> PRT  
 <213> Conus textile  
  
 <220>  
 <221> PEPTIDE  
 <222> (1)..(17)  
 <223> Xaa at residue 14 is Glu or gamma-carboxy Glu; Xaa at residue 7 is  
 s Trp or bromo-Trp; Xaa at residue 17 is Tyr, 125I-Tyr, mono-iodo  
 -Tyr, di-iodo-Tyr, O-sulpho-Tyr or O-phospho-Ty  
  
 <400> 226  
 Phe Cys Cys Asp Ser Asn Xaa Cys His Ile Ser Asp Cys Xaa Cys Cys  
 1 5 10 15

## Xaa

<210> 227  
 <211> 456  
 <212> DNA  
 <213> Conus textile

<220>  
 <221> misc\_feature  
 <222> (1)..(456)  
 <223> n may be any nucleotide

<400> 227  
 caaggaacag tcaacccac agccacgcca agagcagaca gccacagcta cgtgaagaag 60  
 ggtggagaga gggtcgtgat gttgaaaatg ggagtgggtgc tattcatctt cctggtactg 120  
 tttcccctgg caacgctcca gctggatgca gatcaacctg tagaacgata tgcggagaac 180  
 aaacagctcc tcagcccaga tgaaaggagg gaaatcatat tgcattgtct ggggacgcga 240  
 tgctgttctt gggatgtgtg cgaccaccg agttgtactt gctgcggtta gcgccgaaca 300  
 tccatggcgc tgtgctgggc ggttttatcc caacaacgac agcgtttgtt gatttcatgt 360  
 atcattgcgc ccacgtctct tgtctaagaa tgacgaacat gattgcactc tggttcagat 420  
 ttctgtttct tttctgacaa taaatgacaa aacncc 456

<210> 228  
 <211> 70  
 <212> PRT  
 <213> Conus textile

<400> 228  
 Met Leu Lys Met Gly Val Val Leu Phe Ile Phe Leu Val Leu Phe Pro  
 1 5 10 15  
 Leu Ala Thr Leu Gln Leu Asp Ala Asp Gln Pro Val Glu Arg Tyr Ala  
 20 25 30  
 Glu Asn Lys Gln Leu Leu Ser Pro Asp Glu Arg Arg Glu Ile Ile Leu  
 35 40 45  
 His Ala Leu Gly Thr Arg Cys Cys Ser Trp Asp Val Cys Asp His Pro  
 50 55 60  
 Ser Cys Thr Cys Cys Gly  
 65 70

<210> 229  
 <211> 15  
 <212> PRT  
 <213> Conus textile

<220>  
 <221> PEPTIDE  
 <222> (1)..(15)  
 <223> Xaa at residue 10 is Pro or Hyp; Xaa at residue 4 is Trp or bromo  
 -Tr

<400> 229  
 Cys Cys Ser Xaa Asp Val Cys Asp His Xaa Ser Cys Thr Cys Cys  
 1 5 10 15

67

<210> 230  
 <211> 235  
 <212> DNA  
 <213> Conus textile

<400> 230  
 ggatccatga tgtctaaact gggagtcttg ttgaccatct gtctgcttct gtttcccctt 60  
 actgtctctc cgctggatgg agatcaaccc gcagaccaag ctgcagagcg tatgcaggcc 120  
 gagcagcatc ccttgtttga tcagaaaaga cggtgctgca agtttccatg ccccgatagt 180  
 tgcagatatt tgtgttgagg gtgatgataa cgtgttgatg accaactttc tcgag 235

<210> 231  
 <211> 67  
 <212> PRT  
 <213> Conus textile

<400> 231  
 Gly Ser Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu  
 1 5 10 15  
 Leu Phe Pro Leu Thr Ala Leu Pro Leu Asp Gly Asp Gln Pro Ala Asp  
 20 25 30  
 Gln Ala Ala Glu Arg Met Gln Ala Glu Gln His Pro Leu Phe Asp Gln  
 35 40 45  
 Lys Arg Arg Cys Cys Lys Phe Pro Cys Pro Asp Ser Cys Arg Tyr Leu  
 50 55 60

Cys Cys Gly  
 65

<210> 232  
 <211> 16  
 <212> PRT  
 <213> Conus textile

<220>  
 <221> PEPTIDE  
 <222> (1)..(16)  
 <223> Xaa at residue 3 and 8 is Pro or Hyp; Xaa at residue 13 is Tyr, 1  
 25I-Tyr, mono-iodo-Tyr, di-iodo-Tyr, O-sulpho-Tyr or O-phospho-Ty

<400> 232  
 Arg Cys Cys Lys Phe Xaa Cys Xaa Asp Ser Cys Arg Xaa Leu Cys Cys  
 1 5 10 15

<210> 233  
 <211> 321  
 <212> DNA  
 <213> Conus tulipa

<400> 233  
 cgacctcaag agggatcgat agcagttcat gtctaaactg ggagtcttgt tgacaatctg 60  
 tctgcttctg tttcccctta ctgctctgcc gatggatgga gatgaacctg cagaccgacc 120  
 tgcagagcgt atgcaggaca acatttcacg tgagcagcat cccttgtttg aggagagaca 180  
 cggatgttgc aaggggcccg aaggatgctc ctccagagaa tgcagacccc aacattgttg 240

68

cggtcgacga taacgtgttg agggccaact ttgttatcac ggctacgtca agtgtttagt 300

gaataagtaa aatgattgca g 321

<210> 234

<211> 74

<212> PRT

<213> Conus tulipa

<400> 234

Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe Pro  
1 5 10 15

Leu Thr Ala Leu Pro Met Asp Gly Asp Glu Pro Ala Asp Arg Pro Ala  
20 25 30

Glu Arg Met Gln Asp Asn Ile Ser Ser Glu Gln His Pro Leu Phe Glu  
35 40 45

Glu Arg His Gly Cys Cys Lys Gly Pro Glu Gly Cys Ser Ser Arg Glu  
50 55 60

Cys Arg Pro Gln His Cys Cys Gly Arg Arg  
65 70

<210> 235

<211> 21

<212> PRT

<213> Conus tulipa

<220>

<221> PEPTIDE

<222> (1)..(21)

<223> Xaa at residue 8 and 14 is Glu or gamma-carboxy Glu; Xaa at residue 7 and 17 is Pro or Hy

<400> 235

His Gly Cys Cys Lys Gly Xaa Xaa Gly Cys Ser Ser Arg Xaa Cys Arg  
1 5 10 15

Xaa Gln His Cys Cys  
20

<210> 236

<211> 287

<212> DNA

<213> Conus figulinus

<400> 236

caagaaggat cgaatagcagt tcatgatgtc taaactggga gtcttgctga ccatctgtct 60

gcttctgatt ccccttactg ctctttcgct ggatggagat caacctgcag accgacctgc 120

agagcgtatg caggatggaa tttcatctga acagcatccc atgtttgatc ccgtcagacg 180

gtgttgcccg tggccatgca acataggatg cgtaccttgt tgttgatgac cagttttgtt 240

atcgcgccct catcaaatgt ctaatgaata agtaaaacga ttgcagt 287

<210> 237

<211> 67

<212> PRT

<213> Conus figulinus

<400> 237

69

Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Ile  
1 5 10 15

Pro Leu Thr Ala Leu Ser Leu Asp Gly Asp Gln Pro Ala Asp Arg Pro  
20 25 30

Ala Glu Arg Met Gln Asp Gly Ile Ser Ser Glu Gln His Pro Met Phe  
35 40 45

Asp Pro Val Arg Arg Cys Cys Pro Trp Pro Cys Asn Ile Gly Cys Val  
50 55 60

Pro Cys Cys  
65

<210> 238

<211> 14

<212> PRT

<213> Conus figulinus

<220>

<221> PEPTIDE

<222> (1)..(14)

<223> Xaa at residue 3, 5 and 12 is Pro or Hyp; Xaa at residue 4 is Trp  
or bromo-Tr

<400> 238

Cys Cys Xaa Xaa Cys Asn Ile Gly Cys Val Xaa Cys Cys  
1 5 10

<210> 239

<211> 283

<212> DNA

<213> Conus figulinus

<400> 239

caagagggat cgaatagcagt tcatgatgtt taaactggga gtcctgttga ccatctgtat 60  
gcttctgttt ccctttactg ctcttccgct ggatggagag caacctgcag accaaacctgc 120  
agagcgcatg cagtatgaca tggtacgtgc aatgaatccc tggtttgatc ccgtaaaaag 180  
gtgctgctcg aagaactgcg cagtatgcat cccttggtgc ccgtaactga ccagcttgat 240  
tatcgcggcc aaggctctaa tgaataagta aaacgattgc agt 283

<210> 240

<211> 67

<212> PRT

<213> Conus figulinus

<400> 240

Met Met Phe Lys Leu Gly Val Leu Leu Thr Ile Cys Met Leu Leu Phe  
1 5 10 15

Pro Phe Thr Ala Leu Pro Leu Asp Gly Glu Gln Pro Ala Asp Gln Pro  
20 25 30

Ala Glu Arg Met Gln Tyr Asp Met Leu Arg Ala Met Asn Pro Trp Phe  
35 40 45

Asp Pro Val Lys Arg Cys Cys Ser Lys Asn Cys Ala Val Cys Ile Pro  
50 55 60

Cys Cys Pro



65

<210> 241  
 <211> 14  
 <212> PRT  
 <213> Conus figulinus

<220>  
 <221> PEPTIDE  
 <222> (1)..(14)  
 <223> Xaa at residue 11 and 14 is Pro or Hyp

<400> 241  
 Cys Cys Ser Lys Asn Cys Ala Val Cys Ile Xaa Cys Cys Xaa  
 1 5 10

<210> 242  
 <211> 286  
 <212> DNA  
 <213> Conus figulinus

<400> 242  
 caagagggat cgatagcagt tcatgatgtc taaactgaga gtcttgttga ccttatgtct 60  
 gcttctgttt ccccttactg ctcttccgct gaatgaagat caacctgcag agcgtatgca 120  
 ggacgacaat tcatctgagc agcaccctt gtatgaccac aaacgaaagt gttgccggtg 180  
 gccatgcccc gcaagatgcg gctcttgttg cctgtaataa cgtgttggcc aactttgtta 240  
 tcacggccac gtcaaatgtt taatgaataa gtaaaacgat tgcagt 286

<210> 243  
 <211> 64  
 <212> PRT  
 <213> Conus figulinus

<400> 243  
 Met Met Ser Lys Leu Arg Val Leu Leu Thr Leu Cys Leu Leu Leu Phe  
 1 5 10 15

Pro Leu Thr Ala Leu Pro Leu Asn Glu Asp Gln Pro Ala Glu Arg Met  
 20 25 30

Gln Asp Asp Asn Ser Ser Glu Gln His Pro Leu Tyr Asp His Lys Arg  
 35 40 45

Lys Cys Cys Arg Trp Pro Cys Pro Ala Arg Cys Gly Ser Cys Cys Leu  
 50 55 60

<210> 244  
 <211> 15  
 <212> PRT  
 <213> Conus figulinus

<220>  
 <221> PEPTIDE  
 <222> (1)..(15)  
 <223> Xaa at residue 5 and 7 is Pro or Hyp; Xaa at residue 4 is Trp or bromo-Tr

<400> 244  
 Cys Cys Arg Xaa Xaa Cys Xaa Ala Arg Cys Gly Ser Cys Cys Leu  
 1 5 10 15

<210> 245

<211> 301  
 <212> DNA  
 <213> Conus figulinus

<400> 245  
 caagagggat cgatagcagt tcatgatgtc taaactggga gtcttggtga ccttatgtct 60  
 gcttctgttt cccctgactg ctcttccgct ggatgaagat caagctgcag accgacctgc 120  
 agagcgtatg cagggcatgt catctgaaca gcatcccttc ttgatcccg tcaaacggtg 180  
 ttgcgagttg tcacgctgcc ttggatgcgt cccttggtgc acatcttaat aacgtgtgga 240  
 tgaccaactg tgttatcacg gccacgtcaa gtgtctaata aataagtaaa atgattgcag 300  
 t 301

<210> 246  
 <211> 68  
 <212> PRT  
 <213> Conus figulinus

<400> 246  
 Met Met Ser Lys Leu Gly Val Leu Leu Thr Leu Cys Leu Leu Leu Phe  
 1 5 10 15  
 Pro Leu Thr Ala Leu Pro Leu Asp Glu Asp Gln Ala Ala Asp Arg Pro  
 20 25 30  
 Ala Glu Arg Met Gln Gly Met Ser Ser Glu Gln His Pro Phe Phe Asp  
 35 40 45  
 Pro Val Lys Arg Cys Cys Glu Leu Ser Arg Cys Leu Gly Cys Val Pro  
 50 55 60  
 Cys Cys Thr Ser  
 65

<210> 247  
 <211> 16  
 <212> PRT  
 <213> Conus figulinus

<220>  
 <221> PEPTIDE  
 <222> (1)..(16)  
 <223> Xaa at residue 3 and 12 is Pro or Hyp

<400> 247  
 Cys Cys Xaa Leu Ser Arg Cys Leu Gly Cys Val Xaa Cys Cys Thr Ser  
 1 5 10 15

<210> 248  
 <211> 301  
 <212> DNA  
 <213> Conus figulinus

<400> 248  
 caagagggat cgatagcagt tcatgatgtc taaactggga gtcttggtga ccttatgtct 60  
 gcttctgttt cccctgactg ctcttccgct ggatgaagat caacctgcag accgacctgc 120  
 agagcgtatg cagggcatgt catctgaaca gcatcccttc ttgatcccg tcaaacggtg 180  
 ttgcgagttg tcaaaatgcc atggatgcgt cccttggtgc ataccttaat aacgtgcgga 240

72

tgaccaactg tgttatcacg gccacgtcaa gtgtctaatag aataagtaaa atgattgcag 300

t 301

<210> 249

<211> 68

<212> PRT

<213> Conus figulinus

<400> 249

Met Met Ser Lys Leu Gly Val Leu Leu Thr Leu Cys Leu Leu Leu Phe  
1 5 10 15

Pro Leu Thr Ala Leu Pro Leu Asp Glu Asp Gln Pro Ala Asp Arg Pro  
20 25 30

Ala Glu Arg Met Gln Gly Met Ser Ser Glu Gln His Pro Phe Phe Asp  
35 40 45

Pro Val Lys Arg Cys Cys Glu Leu Ser Lys Cys His Gly Cys Val Pro  
50 55 60

Cys Cys Ile Pro  
65

<210> 250

<211> 16

<212> PRT

<213> Conus figulinus

<220>

<221> PEPTIDE

<222> (1)..(16)

<223> Xaa at residue 3 is Glu or gamma-carboxy Glu; Xaa at residue 12 and 16 is Pro or Hy

<400> 250

Cys Cys Xaa Leu Ser Lys Cys His Gly Cys Val Xaa Cys Cys Ile Xaa  
1 5 10 15

<210> 251

<211> 298

<212> DNA

<213> Conus quercinus

<400> 251

caagagggat c gatagcagt tcatgatgtc taaactcgga gtcttggtga ccatctgtct 60

ggttctgttt ccccttacag ctcttcagct ggatggagat caacctgcag accgacctgc 120

agagcgtacg caggacattt catctgaaca gtatcgaaag tttgatcaga gacagaggtg 180

ttgccggtgg ccatgcccg gtagttgcag atgctgccgt tatcggttaac gtgttggtga 240

ccagctttgt tatcacgacc acgccaagtg tctaacgaat aagtaaaatg attgcagt 298

<210> 252

<211> 68

<212> PRT

<213> Conus quercinus

<400> 252

Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Val Leu Phe  
1 5 10 15

Pro Leu Thr Ala Leu Gln Leu Asp Gly Asp Gln Pro Ala Asp Arg Pro

73

20 25 30

Ala Glu Arg Thr Gln Asp Ile Ser Ser Glu Gln Tyr Arg Lys Phe Asp  
35 40 45

Gln Arg Gln Arg Cys Cys Arg Trp Pro Cys Pro Gly Ser Cys Arg Cys  
50 55 60

Cys Arg Tyr Arg  
65

<210> 253  
<211> 18  
<212> PRT  
<213> Conus quercinus

<220>  
<221> PEPTIDE  
<222> (1)..(18)  
<223> Xaa at residue 1 is Gln or pyro-Glu; Xaa at residue 7 and 9 is Pr  
o or Hyp; Xaa at residue 6 is Trp or bromo-Trp; Xaa at residue 17  
is Tyr, 125I-Tyr, mono-iodo-Tyr, di-iodo-Tyr, O-sulpho-Tyr or O-  
phospho-Ty

<400> 253  
Xaa Arg Cys Cys Arg Xaa Xaa Cys Xaa Gly Ser Cys Arg Cys Cys Arg  
1 5 10 15

Xaa Arg

<210> 254  
<211> 313  
<212> DNA  
<213> Conus quercinus

<400> 254  
caagagggat cgatagcagt tcatgatgtc taaactggga gtcttggtga ccatctgtct 60  
gcttctgttt ccccttactg ctcttcact ggatggagat caacctgcag atcaatctgc 120  
agagcgacct gcagagcgta cgcaggacga cattcagcag catccgttat atgatccgaa 180  
aagaaggtgt tgccgttatc catgccccga cagctgccac ggatcttgc tctataagtg 240  
ataacatggt gatggccagc tttgttatca cggccacgtc aagtgtctaa tgaataagta 300  
aaacgattgc agt 313

<210> 255  
<211> 72  
<212> PRT  
<213> Conus quercinus

<400> 255  
Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
1 5 10 15

Pro Leu Thr Ala Leu Pro Leu Asp Gly Asp Gln Pro Ala Asp Gln Ser  
20 25 30

Ala Glu Arg Pro Ala Glu Arg Thr Gln Asp Asp Ile Gln Gln His Pro  
35 40 45

Leu Tyr Asp Pro Lys Arg Arg Cys Cys Arg Tyr Pro Cys Pro Asp Ser  
50 55 60

Cys His Gly Ser Cys Cys Tyr Lys  
65 70

<210> 256  
<211> 18  
<212> PRT  
<213> Conus quercinus

<220>  
<221> PEPTIDE  
<222> (1)..(18)  
<223> Xaa at residue 6 and 8 is Pro or Hyp; Xaa at residue 5 and 17 is Tyr, 125I-Tyr, mono-iodo-Tyr, di-iodo-Tyr, O-sulpho-Tyr or O-phospho-Ty

<400> 256  
Arg Cys Cys Arg Xaa Xaa Cys Xaa Asp Ser Cys His Gly Ser Cys Cys  
1 5 10 15

Xaa Lys

<210> 257  
<211> 256  
<212> DNA  
<213> Conus wittigi

<400> 257  
ggatccatga tgtctaaact gggagtccttg ttgacatct gtctgttct gtttccatt 60  
actgctcttc cgggtgggtgg agatcagcct gcagaccgac ttgcagagcg tatgcaggac 120  
gacacttcat ctgagcagca tccctttgaa aagagactac catcatgttg cgactttgag 180  
aggctttgcg tagtaccagc atgcatacgt catcagtgtt gcacaggata acgtgttgat 240  
gaccaacttt ctcgag 256

<210> 258  
<211> 74  
<212> PRT  
<213> Conus wittigi

<400> 258  
Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
1 5 10 15  
Pro Ile Thr Ala Leu Pro Val Gly Gly Asp Gln Pro Ala Asp Arg Leu  
20 25 30  
Ala Glu Arg Met Gln Asp Asp Thr Ser Ser Glu Gln His Pro Phe Glu  
35 40 45  
Lys Arg Leu Pro Ser Cys Cys Asp Phe Glu Arg Leu Cys Val Val Pro  
50 55 60

Ala Cys Ile Arg His Gln Cys Cys Thr Gly  
65 70

<210> 259  
<211> 23  
<212> PRT  
<213> Conus wittigi

<220>

75

<221> PEPTIDE  
 <222> (1)..(23)  
 <223> Xaa at residue 8 is Glu or gamma-carboxy Glu; Xaa at residue 2 and 14 is Pro or Hy

<400> 259  
 Leu Xaa Ser Cys Cys Asp Phe Xaa Arg Leu Cys Val Val Xaa Ala Cys  
 1 5 10 15

Ile Arg His Gln Cys Cys Thr  
 20

<210> 260  
 <211> 14  
 <212> PRT  
 <213> Conus betulinus

<220>  
 <221> PEPTIDE  
 <222> (1)..(14)  
 <223> Xaa at residue 11 is Pro or Hyp; Xaa at residue 14 is Trp or bromo-Tr

<400> 260  
 Cys Cys Lys Gln Ser Cys Thr Thr Cys Met Xaa Cys Cys Xaa  
 1 5 10

<210> 261  
 <211> 259  
 <212> DNA  
 <213> Conus tulipa

<220>  
 <221> misc\_feature  
 <222> (1)..(259)  
 <223> n may be any nucleotide

<400> 261  
 ggatccatga tgtctaaact gggagtccttg ttgacaatct gtctgcttct gtttcccctt 60  
 actgctctgc cgatggatgg agatgaacct gcagaccgac ctgcagagcg tatgcaggac 120  
 aacatttcac ctgagcagca tcccttggtt gagagagac acggatgttg cgaggggccg 180  
 aagggatgct cctccagaga atgcagaccc caacattgtt gcggtcgacg ataacgtgtt 240  
 gatgaccaac tntctogag 259

<210> 262  
 <211> 75  
 <212> PRT  
 <213> Conus tulipa

<400> 262  
 Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15

Pro Leu Thr Ala Leu Pro Met Asp Gly Asp Glu Pro Ala Asp Arg Pro  
 20 25 30

Ala Glu Arg Met Gln Asp Asn Ile Ser Ser Glu Gln His Pro Leu Phe  
 35 40 45

Glu Glu Arg His Gly Cys Cys Glu Gly Pro Lys Gly Cys Ser Ser Arg  
 50 55 60

76

Glu Cys Arg Pro Gln His Cys Cys Gly Arg Arg  
65 70 75

<210> 263

<211> 21

<212> PRT

<213> Conus tulipa

<220>

<221> PEPTIDE

<222> (1)..(21)

<223> Xaa at residue 5 and 14 is Glu or gamma-carboxy Glu; Xaa at residue 7 and 17 is Pro or Hy

<400> 263

His Gly Cys Cys Xaa Gly Xaa Lys Gly Cys Ser Ser Arg Xaa Cys Arg  
1 5 10 15

Xaa Gln His Cys Cys  
20

<210> 264

<211> 262

<212> DNA

<213> Conus aurisiacus

<220>

<221> misc\_feature

<222> (1)..(262)

<223> n may be any nucleotide

<400> 264

ggatccatga tgtctaaact gggagtottg ttgaccatct gtctacttct gtttccctt 60

actgcttttc cgatggatgg agatcaacct gcagaccaac ctgcagatcg tatgcaggac 120

gacatttcat ctgagcagta tcccttggtt gataagagac aaaagtgttg cactgggagg 180

aaggggtcat gtcctggcaa agcatgcaaa aatctcaaat gttgctctgg acgataacgt 240

gttgatgacc aactttctcg an 262

<210> 265

<211> 76

<212> PRT

<213> Conus aurisiacus

<400> 265

Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
1 5 10 15

Pro Leu Thr Ala Phe Pro Met Asp Gly Asp Gln Pro Ala Asp Gln Pro  
20 25 30

Ala Asp Arg Met Gln Asp Asp Ile Ser Ser Glu Gln Tyr Pro Leu Phe  
35 40 45

Asp Lys Arg Gln Lys Cys Cys Thr Gly Arg Lys Gly Ser Cys Ser Gly  
50 55 60

Lys Ala Cys Lys Asn Leu Lys Cys Cys Ser Gly Arg  
65 70 75

<210> 266

77

<211> 23  
 <212> PRT  
 <213> Conus aurisiacus

<220>  
 <221> PEPTIDE  
 <222> (1)..(23)  
 <223> Xaa at residue 1 is Gln or pyro-Glu

<400> 266  
 Xaa Lys Cys Cys Thr Gly Arg Lys Gly Ser Cys Ser Gly Lys Ala Cys  
 1 5 10 15  
 Lys Asn Leu Lys Cys Cys Ser  
 20

<210> 267  
 <211> 239  
 <212> DNA  
 <213> Conus betulinus

<400> 267  
 ggatccatga tgtctaaact gggagtcttg ttgaccatct gtctgcttct gtttcccctt 60  
 actgctgttc cgttggatgg agatcaacct gcagaccâac ctgcagagcg tatgcagaac 120  
 gaggcagcatc cctcgtttga tcagaaaaga aggtgctgcc ggtggccatg cccaggtata 180  
 tgcgggcatgg ctaggtgttg cttcgtcatg ataacgtgtt gatgaccaac tttctcgag 239

<210> 268  
 <211> 71  
 <212> PRT  
 <213> Conus betulinus

<400> 268  
 Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15  
 Pro Leu Thr Ala Val Pro Leu Asp Gly Asp Gln Pro Ala Asp Gln Pro  
 20 25 30  
 Ala Glu Arg Met Gln Asn Glu Gln His Pro Ser Phe Asp Gln Lys Arg  
 35 40 45  
 Arg Cys Cys Arg Trp Pro Cys Pro Ser Ile Cys Gly Met Ala Arg Cys  
 50 55 60  
 Cys Phe Val Met Ile Thr Cys  
 65 70

<210> 269  
 <211> 23  
 <212> PRT  
 <213> Conus betulinus

<220>  
 <221> PEPTIDE  
 <222> (1)..(23)  
 <223> Xaa at residue 6 and 8 is Pro or Hyp; Xaa at residue 5 is Trp or  
 bromo-Tr

<400> 269  
 Arg Cys Cys Arg Xaa Xaa Cys Xaa Ser Ile Cys Gly Met Ala Arg Cys  
 1 5 10 15



Cys Phe Val Met Ile Thr Cys  
20

<210> 270  
<211> 226  
<212> DNA  
<213> Conus betulinus

<220>  
<221> misc\_feature  
<222> (1)..(226)  
<223> n may be any nucleotide

<400> 270  
ggatccatga tgtctaaact gggagtcttg ttgatcatct gtctgcttct gtttccccctt 60  
actgctgttc cgctggatgg agatcagcct gcagagcgta cgcagatcga gcagcatccc 120  
ttgtttgacc agaaaagaag gtgttgccgg tggccatgcc ccagtagatg cggcatggct 180  
aggtgttgct tcgtcatgat aacgtgttga tgancgacct ctcnag 226

<210> 271  
<211> 67  
<212> PRT  
<213> Conus betulinus

<400> 271  
Met Met Ser Lys Leu Gly Val Leu Leu Ile Ile Cys Leu Leu Leu Phe  
1 5 10 15  
Pro Leu Thr Ala Val Pro Leu Asp Gly Asp Gln Pro Ala Glu Arg Thr  
20 25 30  
Gln Ile Glu Gln His Pro Leu Phe Asp Gln Lys Arg Arg Cys Cys Arg  
35 40 45  
Trp Pro Cys Pro Ser Arg Cys Gly Met Ala Arg Cys Cys Phe Val Met  
50 55 60

Ile Thr Cys  
65

<210> 272  
<211> 23  
<212> PRT  
<213> Conus betulinus

<220>  
<221> PEPTIDE  
<222> (1)..(23)  
<223> Xaa at residue 6 and 8 is Pro or Hyp; Xaa at residue 5 is Trp or  
bromo-Tr

<400> 272  
Arg Cys Cys Arg Xaa Xaa Cys Xaa Ser Arg Cys Gly Met Ala Arg Cys  
1 5 10 15  
Cys Phe Val Met Ile Thr Cys  
20

<210> 273  
<211> 262  
<212> DNA  
<213> Conus parius

<400> 273  
 ggatccatga tgtctaaact gggagtcttg ttgaccatct gtctgcttct gtttcccctt 60  
 actgctcttc cgatggatgg tgatcaacct gcagaccgac ttgtagagcg tatgcaggac 120  
 aacatttcat ctgagcagca tcccttcttt gaaaagagaa gaggaggctg ttgcacacct 180  
 ccgaagaaat gcaaagaccg agcctgcaaa cctgcacgtt gctgcggccc aggataacgt 240  
 gttgatgacc aactttctcg cc 262

<210> 274  
 <211> 76  
 <212> PRT  
 <213> Conus parius

<400> 274  
 Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15  
 Pro Leu Thr Ala Leu Pro Met Asp Gly Asp Gln Pro Ala Asp Arg Leu  
 20 25 30  
 Val Glu Arg Met Gln Asp Asn Ile Ser Ser Glu Gln His Pro Phe Phe  
 35 40 45  
 Glu Lys Arg Arg Gly Gly Cys Cys Thr Pro Pro Lys Lys Cys Lys Asp  
 50 55 60  
 Arg Ala Cys Lys Pro Ala Arg Cys Cys Gly Pro Gly  
 65 70 75

<210> 275  
 <211> 24  
 <212> PRT  
 <213> Conus parius

<220>  
 <221> PEPTIDE  
 <222> (1)..(24)  
 <223> Xaa at residue 7, 8, 18 and 24 is Pro or Hyp

<400> 275  
 Arg Gly Gly Cys Cys Thr Xaa Xaa Lys Lys Cys Lys Asp Arg Ala Cys  
 1 5 10 15  
 Lys Xaa Ala Arg Cys Cys Gly Xaa  
 20

<210> 276  
 <211> 259  
 <212> DNA  
 <213> Conus parius

<400> 276  
 ggatccatga tgtctaaact gggagtcttg ttgaccatct gtctgcttct gtttcccctt 60  
 actgctcttc cgatggatgg tgatcaacct gcagaccgac ttgtagagcg tatgcaggac 120  
 aacatttcat ctgagcagca tcccttcttt gaaaagagaa gagggtgttg cacacctccg 180  
 aggaaatgca aagaccgagc ctgcaaacct gcacgttggt gcggcccagg ataacgtgtt 240  
 gatgaccaac tttctcgag 259

80

<210> 277  
 <211> 75  
 <212> PRT  
 <213> Conus parius

<400> 277  
 Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15  
 Pro Leu Thr Ala Leu Pro Met Asp Gly Asp Gln Pro Ala Asp Arg Leu  
 20 25 30  
 Val Glu Arg Met Gln Asp Asn Ile Ser Ser Glu Gln His Pro Phe Phe  
 35 40 45  
 Glu Lys Arg Arg Gly Cys Cys Thr Pro Pro Arg Lys Cys Lys Asp Arg  
 50 55 60  
 Ala Cys Lys Pro Ala Arg Cys Cys Gly Pro Gly  
 65 70 75

<210> 278  
 <211> 23  
 <212> PRT  
 <213> Conus parius

<220>  
 <221> PEPTIDE  
 <222> (1)..(23)  
 <223> Xaa at residue 6, 7, 17 and 23 is Pro or Hyp

<400> 278  
 Arg Gly Cys Cys Thr Xaa Xaa Arg Lys Cys Lys Asp Arg Ala Cys Lys  
 1 5 10 15  
 Xaa Ala Arg Cys Cys Gly Xaa  
 20

<210> 279  
 <211> 241  
 <212> DNA  
 <213> Conus coronatus

<400> 279  
 ggatccatga tgtctaaact gggagtcttg ttgaccatct gtctgcttct gtttccaatt 60  
 actgcccttc cgctggatga agatcaacct gcagaccgac ctgcagagcg tatgcaggac 120  
 attgcaactg aacagcatcc cttgtttgat cccgtcaaac ggtgctgcga ttggccatgc 180  
 atcccaggat gcaccccttg ttgcttgccct tgataacgtg ttgatgacca actttctoga 240  
 g 241

<210> 280  
 <211> 68  
 <212> PRT  
 <213> Conus coronatus

<400> 280  
 Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15  
 Pro Ile Thr Ala Leu Pro Leu Asp Glu Asp Gln Pro Ala Asp Arg Pro  
 20 25 30

Ala Glu Arg Met Gln Asp Ile Ala Thr Glu Gln His Pro Leu Phe Asp  
 35 40 45

Pro Val Lys Arg Cys Cys Asp Trp Pro Cys Ile Pro Gly Cys Thr Pro  
 50 55 60

Cys Cys Leu Pro  
 65

<210> 281  
 <211> 16  
 <212> PRT  
 <213> Conus coronatus

<220>  
 <221> PEPTIDE  
 <222> (1)..(16)  
 <223> Xaa at residue 5, 8, 12 and 16 is Pro or Hyp; Xaa at residue 4 is  
 Trp or bromo-Trp

<400> 281  
 Cys Cys Asp Xaa Xaa Cys Ile Xaa Gly Cys Thr Xaa Cys Cys Leu Xaa  
 1 5 10 15

<210> 282  
 <211> 244  
 <212> DNA  
 <213> Conus musicus

<400> 282  
 ggatccatga tgtctaaact gggagtcctg ttgaccatct gtctgcttct gtttcctctt 60  
 tctgtctcttc cgatggatga agatcaactt gcagacctac ctgcagagcg tatgcgggac 120  
 actgcaactg tagatcatcc ctccatgat cctgacaaag cgtgctgcga gcagagctgt 180  
 acaacatgct ttccgtgctg ctagccttga acacagtaac gtgttgatga ccaactttct 240  
 cgag 244

<210> 283  
 <211> 65  
 <212> PRT  
 <213> Conus musicus

<400> 283  
 Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15

Pro Leu Ser Ala Leu Pro Met Asp Glu Asp Gln Leu Ala Asp Leu Pro  
 20 25 30

Ala Glu Arg Met Arg Asp Thr Ala Thr Val Asp His Pro Ser Tyr Asp  
 35 40 45

Pro Asp Lys Ala Cys Cys Glu Gln Ser Cys Thr Thr Cys Phe Pro Cys  
 50 55 60

Cys  
 65

<210> 284  
 <211> 14  
 <212> PRT

<213> Conus musicus

<220>

<221> PEPTIDE

<222> (1)..(14)

<223> Xaa at residue 4 is Glu or gamma-carboxy Glu; Xaa at residue 12 is Pro or Hy

<400> 284

Ala Cys Cys Xaa Gln Ser Cys Thr Thr Cys Phe Xaa Cys Cys  
1 5 10

<210> 285

<211> 14

<212> PRT

<213> Conus betulinus

<220>

<221> PEPTIDE

<222> (1)..(14)

<223> Xaa at residue 4 is Glu or gamma-carboxy Glu; Xaa at residue 12 is Pro or Hy

<400> 285

Ala Cys Cys Xaa Gln Ser Cys Thr Thr Cys Met Xaa Cys Cys  
1 5 10

<210> 286

<211> 14

<212> PRT

<213> Conus betulinus

<220>

<221> PEPTIDE

<222> (1)..(14)

<223> Xaa at residue 3 is Glu or gamma-carboxy Glu; Xaa at residue 11 is Pro or Hyp; Xaa at residue 14 is Trp or bromo-Tr

<400> 286

Cys Cys Xaa Gln Ser Cys Thr Thr Cys Met Xaa Cys Cys Xaa  
1 5 10

<210> 287

<211> 235

<212> DNA

<213> Conus pennaceus

<400> 287

ggatccatga tgtctaaact gggagtcttg ttgaccatct gtctgcttct gtttcccctt 60  
actgctcttc cgctggatgg agatcaacct gcataccaag ctgcagagcg tatgcaggcc 120  
gagcatcatc ccttgtttga tcagaaaaga cggtgctgca agtttccatg ccccgatagt 180  
tgcaaattatt tgtgttgccg gtgatgataa catgttgatg accaactttc ttgag 235

<210> 288

<211> 65

<212> PRT

<213> Conus pennaceus

<400> 288

Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
1 5 10 15

83

Pro Leu Thr Ala Leu Pro Leu Asp Gly Asp Gln Pro Ala Tyr Gln Ala  
20 25 30

Ala Glu Arg Met Gln Ala Glu His His Pro Leu Phe Asp Gln Lys Arg  
35 40 45

Arg Cys Cys Lys Phe Pro Cys Pro Asp Ser Cys Lys Tyr Leu Cys Cys  
50 55 60

Gly  
65

<210> 289

<211> 16

<212> PRT

<213> Conus pennaceus

<220>

<221> PEPTIDE

<222> (1)..(16)

<223> Xaa at residue 6 and 8 is Pro or Hyp; Xaa at residue 13 is Tyr, 1  
25I-Tyr, mono-iodo-Tyr, di-iodo-Tyr, O-sulpho-Tyr or O-phospho-Ty

<400> 289

Arg Cys Cys Lys Phe Xaa Cys Xaa Asp Ser Cys Lys Xaa Leu Cys Cys  
1 5 10 15

<210> 290

<211> 241

<212> DNA

<213> Conus pulicarius

<400> 290

ggatccatga tgtctaaact gggagtcttg ttgaccatct gtctgcttct gtttcccctt 60

actgctcttc cgatggatgg tgatcaactt gcagaccgac ttgtagagcg tatgcaggac 120

aacatttcat ctgagcagca tcccttcttt gatcccgta aacgggtgttg cgtcagctgt 180

tacatgggat gcatcccttg ttgcttctag taataacgtg ttgatgacca actttctcga 240

g 241

<210> 291

<211> 67

<212> PRT

<213> Conus pulicarius

<400> 291

Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
1 5 10 15

Pro Leu Thr Ala Leu Pro Met Asp Gly Asp Gln Leu Ala Asp Arg Leu  
20 25 30

Val Glu Arg Met Gln Asp Asn Ile Ser Ser Glu Gln His Pro Phe Phe  
35 40 45

Asp Pro Val Lys Arg Cys Cys Val Ser Cys Tyr Met Gly Cys Ile Pro  
50 55 60

Cys Cys Phe  
65

<210> 292  
 <211> 14  
 <212> PRT  
 <213> Conus pulicarius

<220>  
 <221> PEPTIDE  
 <222> (1)..(14)  
 <223> Xaa at residue 11 is Pro or Hyp; Xaa at residue 6 is Tyr, 125I-Tyr, mono-iodo-Tyr, di-iodo-Tyr, O-sulpho-Tyr or O-phospho-Ty

<400> 292  
 Cys Cys Val Ser Cys Xaa Met Gly Cys Ile Xaa Cys Cys Phe  
 1 5 10

<210> 293  
 <211> 244  
 <212> DNA  
 <213> Conus pulicarius

<400> 293  
 ggatccatga tgtctaaact gggagtcttg ttgaccgtct gtctgcttct gtgtcccctt 60  
 actgctcttc cactggatga agatcaactt gcagaccgac ctgcagagcg tatgcaggat 120  
 gacacttcag ctgcacagat tttcgggttt gatcccgta aacggtgctg caaattgcta 180  
 tgctactcgg gatgcactcc ttgttgccat atttgataac gtgttgatga ccaactttct 240  
 cgag 244

<210> 294  
 <211> 67  
 <212> PRT  
 <213> Conus pulicarius

<400> 294  
 Met Met Ser Lys Leu Gly Val Leu Leu Thr Val Cys Leu Leu Leu Cys  
 1 5 10 15

Pro Leu Thr Ala Leu Pro Leu Asp Glu Asp Gln Leu Ala Asp Arg Pro  
 20 25 30

Ala Glu Arg Met Gln Asp Asp Thr Ser Ala Ala Gln Ile Phe Gly Phe  
 35 40 45

Asp Pro Val Lys Arg Cys Cys Lys Leu Leu Cys Gly Cys Thr Pro Cys  
 50 55 60

Cys His Ile  
 65

<210> 295  
 <211> 16  
 <212> PRT  
 <213> Conus pulicarius

<220>  
 <221> PEPTIDE  
 <222> (1)..(16)  
 <223> Xaa at residue 12 is Pro or Hyp; Xaa at residue 7 is Tyr, 125I-Tyr, mono-iodo-Tyr, di-iodo-Tyr, O-sulpho-Tyr or O-phospho-Ty

<400> 295  
 Cys Cys Lys Leu Leu Cys Xaa Ser Gly Cys Thr Xaa Cys Cys His Ile

85

1 5 10 15

<210> 296  
 <211> 259  
 <212> DNA  
 <213> Conus rattus

<400> 296  
 ggatccatga tgtctaaact gggagtcttg ttgaccatct gtctgcttgt gtttccgctt 60  
 actgctcttc cgatggatgg tgatcaacct gcagaccgac ttgtagagcg tatacaggac 120  
 aacatttcat ctgagcagca tcccttcttt gaaaagagaa gaggetgttg cgcacctccg 180  
 aggaaatgca aagaccgagc ctgcaaacct gcacgttgct gcggcccagg ataacgtgtt 240  
 gatgaccaac tttctcgag 259

<210> 297  
 <211> 75  
 <212> PRT  
 <213> Conus rattus

<400> 297  
 Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Val Phe  
 1 5 10 15  
 Pro Leu Thr Ala Leu Pro Met Asp Gly Asp Gln Pro Ala Asp Arg Leu  
 20 25 30  
 Val Glu Arg Ile Gln Asp Asn Ile Ser Ser Glu Gln His Pro Phe Phe  
 35 40 45  
 Glu Lys Arg Arg Gly Cys Cys Ala Pro Pro Arg Lys Cys Lys Asp Arg  
 50 55 60  
 Ala Cys Lys Pro Ala Arg Cys Cys Gly Pro Gly  
 65 70 75

<210> 298  
 <211> 23  
 <212> PRT  
 <213> Conus rattus

<220>  
 <221> PEPTIDE  
 <222> (1)..(23)  
 <223> Xaa at residue 6, 7, 17 and 23 is Pro or Hyp

<400> 298  
 Arg Gly Cys Cys Ala Xaa Xaa Arg Lys Cys Lys Asp Arg Ala Cys Lys  
 1 5 10 15  
 Xaa Ala Arg Cys Cys Gly Xaa  
 20

<210> 299  
 <211> 262  
 <212> DNA  
 <213> Conus stercusmuscarum

<400> 299  
 ggatccatga tgtctaaact gggagtcttg ttgacaatct gtctgcttct gtttcccctt 60  
 attgctcttc cgctggatgg agatcaacct gcagaccgac ctgcagagcg tatgcaggac 120



86

gacatttcat ctgagaagca tcccttggtt gataagagac aacggtgttg caatgggcgg 180  
 aggggatgct ccagcagatg gtgcagagat cactcacgtt gttgcggtcg acgataacgt 240  
 gttgatgacc aactttctcg ag 262

<210> 300  
 <211> 76  
 <212> PRT  
 <213> Conus stercusmuscarum

<400> 300  
 Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15  
 Pro Leu Ile Ala Leu Pro Leu Asp Gly Asp Gln Pro Ala Asp Arg Pro  
 20 25 30  
 Ala Glu Arg Met Gln Asp Asp Ile Ser Ser Glu Lys His Pro Leu Phe  
 35 40 45  
 Asp Lys Arg Gln Arg Cys Cys Asn Gly Arg Arg Gly Cys Ser Ser Arg  
 50 55 60  
 Trp Cys Arg Asp His Ser Arg Cys Cys Gly Arg Arg  
 65 70 75

<210> 301  
 <211> 22  
 <212> PRT  
 <213> Conus stercusmuscarum

<220>  
 <221> PEPTIDE  
 <222> (1)..(22)  
 <223> Xaa at residue 1 is Gln or pyro-Glu; Xaa at residue 14 is Trp or  
 bromo-Tr

<400> 301  
 Xaa Arg Cys Cys Asn Gly Arg Arg Gly Cys Ser Ser Arg Xaa Cys Arg  
 1 5 10 15  
 Asp His Ser Arg Cys Cys  
 20

<210> 302  
 <211> 241  
 <212> DNA  
 <213> Conus ebraceus

<400> 302  
 ggatccatga tgtctaaact gggagtcttg ttgaccatct gtctgcttct gtttccctt 60  
 actgctcttc cactggatga aggtcaacct gcagacctac ctgcagagcg tatgcaggac 120  
 attgcaactg aacagcatcc cttgtttgat cctgtcaaac ggtgttgcca gcagccatgc 180  
 tacatgggat gcaccccttg ttgcttctaa taataacgtg ttgatgacca aactttctcga 240  
 g 241

<210> 303  
 <211> 67  
 <212> PRT

87

&lt;213&gt; Conus ebraceus

&lt;400&gt; 303

Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15

Pro Leu Thr Ala Leu Pro Leu Asp Glu Gly Gln Pro Ala Asp Leu Pro  
 20 25 30

Ala Glu Arg Met Gln Asp Ile Ala Thr Glu Gln His Pro Leu Phe Asp  
 35 40 45

Pro Val Lys Arg Cys Cys Glu Gln Pro Cys Tyr Met Gly Cys Ile Pro  
 50 55 60

Cys Cys Phe  
 65

&lt;210&gt; 304

&lt;211&gt; 15

&lt;212&gt; PRT

&lt;213&gt; Conus ebraceus

&lt;220&gt;

&lt;221&gt; PEPTIDE

&lt;222&gt; (1)..(15)

<223> Xaa at residue 3 is Glu or gamma-carboxy Glu; Xaa at residue 5 and  
 12 is Pro or Hyp; Xaa at residue 7 is Tyr, 125I-Tyr, mono-iodo-  
 Tyr, di-iodo-Tyr, O-sulpho-Tyr or O-phospho-Ty

&lt;400&gt; 304

Cys Cys Xaa Gln Xaa Cys Xaa Met Gly Cys Ile Xaa Cys Cys Phe  
 1 5 10 15

&lt;210&gt; 305

&lt;211&gt; 241

&lt;212&gt; DNA

&lt;213&gt; Conus ebraceus

&lt;400&gt; 305

ggatccatga tgtctaaact gggagtccttg ttgacccatct gtcctgcttct gtttccctt 60

actgctcttc cactggatga agatcaacct gcagacctac ctgcagagcg tatgcaggac 120

attgcaactg aacagcatcc cttgtttgat cctgtcaaac ggtgctgctgc gcagccatgc 180

tacatgggat gcatcccttg ttgcttctaa taataacgtg ttgatgacca actttctoga 240

g 241

&lt;210&gt; 306

&lt;211&gt; 67

&lt;212&gt; PRT

&lt;213&gt; Conus ebraceus

&lt;400&gt; 306

Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15

Pro Leu Thr Ala Leu Pro Leu Asp Glu Asp Gln Pro Ala Asp Leu Pro  
 20 25 30

Ala Glu Arg Met Gln Asp Ile Ala Thr Glu Gln His Pro Leu Phe Asp  
 35 40 45

Pro Val Lys Arg Cys Cys Ala Gln Pro Cys Tyr Met Gly Cys Ile Pro

88

50 55 60

Cys Cys Phe  
65

<210> 307  
<211> 15  
<212> PRT  
<213> Conus ebraceus

<220>  
<221> PEPTIDE  
<222> (1)..(15)  
<223> Xaa at residue 5 and 12 is Pro or Hyp; Xaa at residue 7 is Tyr, 1  
25I-Tyr, mono-iodo-Tyr, di-iodo-Tyr, O-sulpho-Tyr or O-phospho-Ty

<400> 307  
Cys Cys Ala Gln Xaa Cys Xaa Met Gly Cys Ile Xaa Cys Cys Phe  
1 5 10 15

<210> 308  
<211> 238  
<212> DNA  
<213> Conus flavidus

<400> 308  
ggatccatga tgtctaaact gggagtcttg ttgaccatct gtctgcttct gtttcccctt 60  
actgctgttc cggtggatgg agatcaacct gcagaccagc ctgcagagcg tatgcagaac 120  
gagcagcatc ccttgtttga tcagaaaaga aggtgctgcc ggtggccatg cccaggtata 180  
tgcggcgatgg ctagggtgttg ctgcgtcatga taacgtgttg atgaccaact ttctcgag 238

<210> 309  
<211> 67  
<212> PRT  
<213> Conus flavidus

<400> 309  
Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
1 5 10 15  
Pro Leu Thr Ala Val Pro Leu Asp Gly Asp Gln Pro Ala Asp Gln Pro  
20 25 30  
Ala Glu Arg Met Gln Asn Glu Gln His Pro Leu Phe Asp Gln Lys Arg  
35 40 45  
Arg Cys Cys Arg Trp Pro Cys Pro Ser Ile Cys Gly Met Ala Arg Cys  
50 55 60

Cys Ser Ser  
65

<210> 310  
<211> 19  
<212> PRT  
<213> Conus flavidus

<220>  
<221> PEPTIDE  
<222> (1)..(19)  
<223> Xaa at residue 6 and 8 is Pro or Hyp; Xaa at residue 5 is Trp or

bromo-Tr

&lt;400&gt; 310

Arg Cys Cys Arg Xaa Xaa Cys Xaa Ser Ile Cys Gly Met Ala Arg Cys  
 1 5 10 15

Cys Ser Ser

&lt;210&gt; 311

&lt;211&gt; 245

&lt;212&gt; DNA

&lt;213&gt; Conus miliaris

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; (1)..(245)

&lt;223&gt; n may be any nucleotide

&lt;400&gt; 311

ggatccatga tgtctaaact gggagtcttg ttgaccatct gtctgcttct gtttccaatt 60  
 actgcccttc cactggatga agatcaacct gcagaccgac ctgcagagcg tatgcaggac 120  
 attgcaactg aacagcatcc ctgtttgat cccgtcaaac ggtgttgca ttggccatgc 180  
 agcgcaggat gctacccttg ttgcttcct taataacgtg ttgatgacca actnangnaa 240  
 aaaaaa 245

&lt;210&gt; 312

&lt;211&gt; 68

&lt;212&gt; PRT

&lt;213&gt; Conus miliaris

&lt;400&gt; 312

Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15

Pro Ile Thr Ala Leu Pro Leu Asp Glu Asp Gln Pro Ala Asp Arg Pro  
 20 25 30

Ala Glu Arg Met Gln Asp Ile Ala Thr Glu Gln His Pro Leu Phe Asp  
 35 40 45

Pro Val Lys Arg Cys Cys Asp Trp Pro Cys Ser Ala Gly Cys Tyr Pro  
 50 55 60

Cys Cys Phe Pro  
 65

&lt;210&gt; 313

&lt;211&gt; 16

&lt;212&gt; PRT

&lt;213&gt; Conus miliaris

&lt;220&gt;

&lt;221&gt; PEPTIDE

&lt;222&gt; (1)..(16)

<223> Xaa at residue 5, 12 and 16 is Pro or Hyp; Xaa at residue 4 is Trp or bromo-Trp; Xaa at residue 11 is Tyr, 125I-Tyr, mono-iodo-Tyr, di-iodo-Tyr, O-sulpho-Tyr or O-phospho-Ty

&lt;400&gt; 313

Cys Cys Asp Xaa Xaa Cys Ser Ala Gly Cys Xaa Xaa Cys Cys Phe Xaa  
 1 5 10 15

<210> 314  
 <211> 230  
 <212> DNA  
 <213> Conus miliaris

<220>  
 <221> misc\_feature  
 <222> (1)..(230)  
 <223> n may be any nucleotide  
 <400> 314  
 ggatccatga tgtctaaact gggagtgggtg ccattcgtct ttctggtcct gtttccctg 60  
 gcaacactcc aactggatgc agatcaacct gcagaccgac ctgcgcgtaa aaagggcatt 120  
 gcaactaaac ggcattccctt gtctgatact gtcagagggt gttgccctcc aatgtgcaca 180  
 ccatgcttcc cttgctgttt tcgttaataa cgtgttgatg natgatgnan 230

<210> 315  
 <211> 66  
 <212> PRT  
 <213> Conus miliaris

<400> 315  
 Met Met Ser Lys Leu Gly Val Val Pro Phe Val Phe Leu Val Leu Phe  
 1 5 10 15  
 Pro Leu Ala Thr Leu Gln Leu Asp Ala Asp Gln Pro Ala Asp Arg Pro  
 20 25 30  
 Ala Arg Lys Lys Gly Ile Ala Thr Lys Arg His Pro Leu Ser Asp Pro  
 35 40 45  
 Val Arg Gly Cys Cys Pro Pro Met Cys Thr Pro Cys Phe Pro Cys Cys  
 50 55 60  
 Phe Arg  
 65

<210> 316  
 <211> 16  
 <212> PRT  
 <213> Conus miliaris

<220>  
 <221> PEPTIDE  
 <222> (1)..(16)  
 <223> Xaa at residue 4, 9 and 12 is Pro or Hyp; Xaa at residue 5 is Tyr  
 , 125I-Tyr, mono-iodo-Tyr, di-iodo-Tyr, O-sulpho-Tyr or O-phospho  
 -Ty

<400> 316  
 Gly Cys Cys Xaa Xaa Met Cys Thr Xaa Cys Phe Xaa Cys Cys Phe Arg  
 1 5 10 15

<210> 317  
 <211> 295  
 <212> DNA  
 <213> Conus ammiralis

<400> 317  
 caagagggat cgatagcagt tcatgatgtc taaactggga gtcttggtga ccatctgtct 60  
 gcttctgttt ccccttactg ctcttccgct ggatggagat caacctgcag accaagctgc 120

agagcgtatg caggccgagc agcatccctt gtttgatcag aaaagacggt gttgcaggtt 180  
 tccatgcccc gatacttgca gacatttggtg ttgcgggtga tgataacgtg ctgatgaccc 240  
 actttgtcat cacggctacg tcaagtgtct aatgaataag taaaatgatt gcagt 295  
 <210> 318  
 <211> 65  
 <212> PRT  
 <213> Conus ammiralis  
  
 <400> 318  
 Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15  
 Pro Leu Thr Ala Leu Pro Leu Asp Gly Asp Gln Pro Ala Asp Gln Ala  
 20 25 30  
 Ala Glu Arg Met Gln Ala Glu Gln His Pro Leu Phe Asp Gln Lys Arg  
 35 40 45  
 Arg Cys Cys Arg Phe Pro Cys Pro Asp Thr Cys Arg His Leu Cys Cys  
 50 55 60  
 Gly  
 65  
  
 <210> 319  
 <211> 16  
 <212> PRT  
 <213> Conus ammiralis  
  
 <220>  
 <221> PEPTIDE  
 <222> (1)..(16)  
 <223> Xaa at residue 6 and 8 is Pro or Hyp  
  
 <400> 319  
 Arg Cys Cys Arg Phe Xaa Cys Xaa Asp Thr Cys Arg His Leu Cys Cys  
 1 5 10 15  
  
 <210> 320  
 <211> 267  
 <212> DNA  
 <213> Conus ammiralis  
  
 <400> 320  
 caagagggat cgatagcagt tcatgatgtt taaactggga gtcttgctga ccatctgtct 60  
 acttctgttt tcccttaatg ctgttccgct ggatggagat caacctgcag accaacctgc 120  
 agagcgtctg ctggacgaca ttcatctga aaataatccc ttttatgac ccgccaaacg 180  
 gtgttgcatg acttgcttcg gttgcacacc ttgttggtga tgaccagcct catcaagtgt 240  
 ctaacgaata agtaaaacga ttgcagt 267  
  
 <210> 321  
 <211> 66  
 <212> PRT  
 <213> Conus ammiralis  
  
 <400> 321  
 Met Met Phe Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15

Ser Leu Asn Ala Val Pro Leu Asp Gly Asp Gln Pro Ala Asp Gln Pro  
20 25 30

Ala Glu Arg Leu Leu Asp Asp Ile Ser Ser Glu Asn Asn Pro Phe Tyr  
35 40 45

Asp Pro Ala Lys Arg Cys Cys Met Thr Cys Phe Gly Cys Thr Pro Cys  
50 55 60

Cys Gly  
65

<210> 322

<211> 12

<212> PRT

<213> Conus ammiralis

<220>

<221> PEPTIDE

<222> (1)..(12)

<223> Xaa at residue 10 is Pro or Hyp

<400> 322

Cys Cys Met Thr Cys Phe Gly Cys Thr Xaa Cys Cys  
1 5 10

<210> 323

<211> 294

<212> DNA

<213> Conus ammiralis

<400> 323

caagaaggat cgatagcagt tcatgatgtc taaactggga gccttggtga ccatctgtct 60

acttctgttt tcccttactg ctgttccgct ggatggagat caacatgcag accaacctgc 120

agagcgtctg caggaccgcc ttccaactga aaatcatccc ttatatgata ccgtaaaccg 180

gtgttgcat gatctggaat ggcactattc ttgctggcct tgctgtattt tttcataacc 240

tttgttatcg cggcctcatc ctagtgtcaa atgaataagt aaaacgattg cagt 294

<210> 324

<211> 71

<212> PRT

<213> Conus ammiralis

<400> 324

Met Met Ser Lys Leu Gly Ala Leu Leu Thr Ile Cys Leu Leu Leu Phe  
1 5 10 15

Ser Leu Thr Ala Val Pro Leu Asp Gly Asp Gln His Ala Asp Gln Pro  
20 25 30

Ala Glu Arg Leu Gln Asp Arg Leu Pro Thr Glu Asn His Pro Leu Tyr  
35 40 45

Asp Pro Val Lys Arg Cys Cys Asp Asp Ser Glu Cys Asp Tyr Ser Cys  
50 55 60

Trp Pro Cys Cys Ile Phe Ser  
65 70

<210> 325

<211> 18

<212> PRT  
<213> Conus ammiralis

<220>  
<221> PEPTIDE  
<222> (1)..(18)  
<223> Xaa at residue 6 is Glu or gamma-carboxy Glu; Xaa at residue 13 is Pro or Hyp; Xaa at residue 12 is Trp or bromo-Trp; Xaa at residue 9 is Tyr, 125I-Tyr, mono-iodo-Tyr, di-iodo-Tyr, O-sulpho-Tyr or O-phospho-Ty

<400> 325  
Cys Cys Asp Asp Ser Xaa Cys Asp Xaa Ser Cys Xaa Xaa Cys Cys Ile  
1 5 10 15

Phe Ser

<210> 326  
<211> 284  
<212> DNA  
<213> Conus ammiralis

<400> 326  
caagagggat cgatagcagt tcatgatgtt taaactcgga gtcttgctga ccattctgtct 60  
acttctgttt tccctaattg ctgttccgct ggatggagat caacatgcag accaacctgc 120  
agagcgtctg caggaccgcc ttccaactga aaatcatccc ttatatgatc ccgtcaaacg 180  
gtgttgaggg ttgttatgcc tcagttgcaa cccttggtgt ggatgaccag ctttgttattc 240  
acggcctcat caagtgtcta atgaataagt aaaacgattg cagt 284

<210> 327  
<211> 67  
<212> PRT  
<213> Conus ammiralis

<400> 327  
Met Met Phe Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
1 5 10 15

Ser Leu Ile Ala Val Pro Leu Asp Gly Asp Gln His Ala Asp Gln Pro  
20 25 30

Ala Glu Arg Leu Gln Asp Arg Leu Pro Thr Glu Asn His Pro Leu Tyr  
35 40 45

Asp Pro Val Lys Arg Cys Cys Arg Leu Leu Cys Leu Ser Cys Asn Pro  
50 55 60

Cys Cys Gly  
65

<210> 328  
<211> 13  
<212> PRT  
<213> Conus ammiralis

<220>  
<221> PEPTIDE  
<222> (1)..(13)  
<223> Xaa at residue 11 is Pro or Hyp

<400> 328



94

Cys Cys Arg Leu Leu Cys Leu Ser Cys Asn Xaa Cys Cys  
 1 5 10

<210> 329  
 <211> 289  
 <212> DNA  
 <213> Conus ammiralis

<400> 329  
 caagaaggat cgatagcagt tcatgatgtc taaactggga gccttggtga ccatctgtct 60  
 acttctgttt tccottactg ctgttccgct ggatggagat caacatgcag accaacctgc 120  
 agagcgtctg caggaccgca ttccaactga agatcatccc ttattigatc ccaacaaacg 180  
 gtgttgcatg gattcggaat gcggctattc atgtggcct tgctgttatg gataagcttt 240  
 gttatcgcg cctcatccag tgtcaacgaa taagtaaaac gattgcagt 289

<210> 330  
 <211> 70  
 <212> PRT  
 <213> Conus ammiralis

<400> 330  
 Met Met Ser Lys Leu Gly Ala Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15  
 Ser Leu Thr Ala Val Pro Leu Asp Gly Asp Gln His Ala Asp Gln Pro  
 20 25 30  
 Ala Glu Arg Leu Gln Asp Arg Ile Pro Thr Glu Asp His Pro Leu Phe  
 35 40 45  
 Asp Pro Asn Lys Arg Cys Cys Asp Asp Ser Glu Cys Gly Tyr Ser Cys  
 50 55 60

Trp Pro Cys Cys Tyr Gly  
 65 70

<210> 331  
 <211> 16  
 <212> PRT  
 <213> Conus ammiralis

<220>  
 <221> PEPTIDE  
 <222> (1)..(16)  
 <223> Xaa at residue6 is Glu or gamma-carboxy Glu; Xaa at residue 13 is Pro or Hyp; Xaa at residue 12 is Trp or bromo-Trp; Xaa at residue 9 and 16 is Tyr, 125I-Tyr, mono-iodo-Tyr, di-iodo-Tyr, O-sulpho-Tyr or O-phospho-Ty

<400> 331  
 Cys Cys Asp Asp Ser Xaa Cys Gly Xaa Ser Cys Xaa Xaa Cys Cys Xaa  
 1 5 10 15

<210> 332  
 <211> 272  
 <212> DNA  
 <213> Conus spurius

<400> 332  
 caagaaggat cgatagcagt tcatgatgtc taaactggga gtcttgctga ccatctgtct 60

95

gcttctgttt ccacgtactt ctcttccgct ggatggagat caacctgcag tccgatctgc 120  
 aaagcgtatg cattcatcta tacagcgctg tttctttgat cccgtcaaac ggtgttgccc 180  
 tagatgcagc gagtgaacc cttgttgtgg atgaccagct ttgtcatcgc ggcctcatta 240  
 agtgtctaata gaataagtaa aatgattgca gt 272

<210> 333  
 <211> 63  
 <212> PRT  
 <213> Conus spurius

<400> 333  
 Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15  
 Pro Arg Thr Ser Leu Pro Leu Asp Gly Asp Gln Pro Ala Val Arg Ser  
 20 25 30  
 Ala Lys Arg Met His Ser Ser Ile Gln Arg Arg Phe Phe Asp Pro Val  
 35 40 45  
 Lys Arg Cys Cys Pro Arg Cys Ser Glu Cys Asn Pro Cys Cys Gly  
 50 55 60

<210> 334  
 <211> 12  
 <212> PRT  
 <213> Conus spurius

<220>  
 <221> PEPTIDE  
 <222> (1)..(12)  
 <223> Xaa at residue 7 is Glu or gamma-carboxy Glu; Xaa at residue 3 and 10 is Pro or Hy

<400> 334  
 Cys Cys Xaa Arg Cys Ser Xaa Cys Asn Xaa Cys Cys  
 1 5 10

<210> 335  
 <211> 293  
 <212> DNA  
 <213> Conus omaria

<400> 335  
 caagagggat cgatagcagt tcatgatgtc taaactggga gtctcgttga ccatctgtct 60  
 acttctattt tccottactg ctgttccgct tgatggagat caacatgcag accaacctgc 120  
 agagcgtctg cagggcgaca ttttatctga aaagcatccc ttatttaatc ccgtaaaccg 180  
 gtgttgcatg gaggaagaat gcagcagtcg atgctggcct tgtgtgtggg ggtgatcagc 240  
 tttgttatcg cggcctcatc aagtgtctaa tgaataagta aatgattgc agt 293

<210> 336  
 <211> 70  
 <212> PRT  
 <213> Conus omaria

<400> 336  
 Met Met Ser Lys Leu Gly Val Ser Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15

Ser Leu Thr Ala Val Pro Leu Asp Gly Asp Gln His Ala Asp Gln Pro  
20 25 30

Ala Glu Arg Leu Gln Gly Asp Ile Leu Ser Glu Lys His Pro Leu Phe  
35 40 45

Asn Pro Val Lys Arg Cys Cys Asp Glu Glu Glu Cys Ser Ser Ala Cys  
50 55 60

Trp Pro Cys Cys Trp Gly  
65 70

<210> 337

<211> 16

<212> PRT

<213> Conus omaria

<220>

<221> PEPTIDE

<222> (1)..(16)

<223> Xaa at residue 4, 5 and 6 is Glu or gamma-carboxy Glu; Xaa at residue 13 is Pro or Hyp; Xaa at residue 12 and 16 is Trp or bromo-Tr

<400> 337

Cys Cys Asp Xaa Xaa Xaa Cys Ser Ser Ala Cys Xaa Xaa Cys Cys Xaa  
1 5 10 15

<210> 338

<211> 293

<212> DNA

<213> Conus omaria

<400> 338

caagaaggat cgatagcagt tcatgatgtc taaactggga gtcttggtga tcattctgtct 60

acttctgtgt ccccttactg ctgtttctgga ggatggagat caacctgcag accgacctgc 120

agagcgtatg caggacgaca ttccaactga gcatcatccc ttttatgatc ccgtcaaacg 180

gtgttgcaag tacgggtgga catgcttgct aggatgcact ccttggtgatt gttgaccagt 240

tttgttatcg cggcctcgtc aagtgtctaa tgaataagta aaacgattgc agt 293

<210> 339

<211> 70

<212> PRT

<213> Conus omaria

<400> 339

Met Met Ser Lys Leu Gly Val Leu Leu Ile Ile Cys Leu Leu Leu Cys  
1 5 10 15

Pro Leu Thr Ala Val Leu Glu Asp Gly Asp Gln Pro Ala Asp Arg Pro  
20 25 30

Ala Glu Arg Met Gln Asp Asp Ile Ser Thr Glu His His Pro Phe Tyr  
35 40 45

Asp Pro Val Lys Arg Cys Cys Lys Tyr Gly Trp Thr Cys Leu Leu Gly  
50 55 60

Cys Thr Pro Cys Asp Cys  
65 70

<210> 340  
 <211> 17  
 <212> PRT  
 <213> Conus omaria

<220>  
 <221> PEPTIDE  
 <222> (1)..(17)  
 <223> Xaa at residue is 14 Pro or Hyp; Xaa at residue 6 is Trp or bromo-Trp; Xaa at residue 4 is Tyr, 125I-Tyr, mono-iodo-Tyr, di-iodo-Tyr, O-sulpho-Tyr or O-phospho-Ty

<400> 340  
 Cys Cys Lys Xaa Gly Xaa Thr Cys Leu Leu Gly Cys Thr Xaa Cys Asp  
 1 5 10 15

Cys

<210> 341  
 <211> 290  
 <212> DNA  
 <213> Conus omaria

<400> 341  
 caagagggat cgatagcagt tcatgatgtc tatactggga gtcttggtga tcatctgtct 60  
 acttctgtgt ccccttactg ctgttctgga ggatggagat caacctgcag accgacctgc 120  
 agagcgtatg caggacggca ttcatctga acatcatccc tttttggatc ccgtcaaacg 180  
 gtgttgccat ctattggcat gccgctttgg atgctcgctt tgttggtggt gaccagcttt 240  
 gttatcgcg cctcatcaag tgtctaataa ataagtaaaa cgattgcagt 290

<210> 342  
 <211> 69  
 <212> PRT  
 <213> Conus omaria

<400> 342  
 Met Met Ser Ile Leu Gly Val Leu Leu Ile Ile Cys Leu Leu Leu Cys  
 1 5 10 15  
 Pro Leu Thr Ala Val Leu Glu Asp Gly Asp Gln Pro Ala Asp Arg Pro  
 20 25 30  
 Ala Glu Arg Met Gln Asp Gly Ile Ser Ser Glu His His Pro Phe Leu  
 35 40 45  
 Asp Pro Val Lys Arg Cys Cys His Leu Leu Ala Cys Arg Phe Gly Cys  
 50 55 60

Ser Pro Cys Cys Trp  
 65

<210> 343  
 <211> 16  
 <212> PRT  
 <213> Conus omaria

<220>  
 <221> PEPTIDE  
 <222> (1)..(16)  
 <223> Xaa at residue 13 is Pro or Hyp; Xaa at residue 16 is Trp or bromo-Trp

&lt;400&gt; 343

Cys Cys His Leu Leu Ala Cys Arg Phe Gly Cys Ser Xaa Cys Cys Xaa  
 1 5 10 15

&lt;210&gt; 344

&lt;211&gt; 293

&lt;212&gt; DNA

&lt;213&gt; Conus omaria

&lt;400&gt; 344

caagaaggat cgatagcagt tcatgatgtc taaactggga gtcttggtga tcatctgtct 60  
 acttctttgt ccccttactg ctgttccgca ggatggagat caacctgcag accgacctgc 120  
 agagcgtatg caggggcgca tttcatctga acatcatccc ttttttgatc ccgtcaaacg 180  
 gtgttgacagg tacgggtgga catgctggct aggatgcact ccctgtgggt gttgaccagc 240  
 tttgttatcg cggcctcatc aagtgtctaa tgaataagta aaacgattgc agt 293

&lt;210&gt; 345

&lt;211&gt; 70

&lt;212&gt; PRT

&lt;213&gt; Conus omaria

&lt;400&gt; 345

Met Met Ser Lys Leu Gly Val Leu Leu Ile Ile Cys Leu Leu Leu Cys  
 1 5 10 15

Pro Leu Thr Ala Val Pro Gln Asp Gly Asp Gln Pro Ala Asp Arg Pro  
 20 25 30

Ala Glu Arg Met Gln Gly Gly Ile Ser Ser Glu His His Pro Phe Phe  
 35 40 45

Asp Pro Val Lys Arg Cys Cys Arg Tyr Gly Trp Thr Cys Trp Leu Gly  
 50 55 60

Cys Thr Pro Cys Gly Cys  
 65 70

&lt;210&gt; 346

&lt;211&gt; 17

&lt;212&gt; PRT

&lt;213&gt; Conus omaria

&lt;220&gt;

&lt;221&gt; PEPTIDE

&lt;222&gt; (1)..(17)

<223> Xaa at residue 14 is Pro or Hyp; Xaa at residue 6 and 9 is Trp or  
 bromo-Trp; Xaa at residue 4 is Tyr, 125I-Tyr, mono-iodo-Tyr, di-  
 iodo-Tyr, O-sulpho-Tyr or O-phospho-Ty

&lt;400&gt; 346

Cys Cys Arg Xaa Gly Xaa Thr Cys Xaa Leu Gly Cys Thr Xaa Cys Gly  
 1 5 10 15

Cys

&lt;210&gt; 347

&lt;211&gt; 293

&lt;212&gt; DNA

&lt;213&gt; Conus episcopatus

<400> 347  
 caagaaggat cgatagcagt tcatgatgtc taaactggga gtcttggtga ccattctgtct 60  
 acttctgttt tcccttattg ctgttccgct tgatggagat caacatgcag accaacctgc 120  
 agagcgtctg cagggcgaca ttttatctga aaagcatccc ttatttatgc ctgtcaaacg 180  
 gtgttgcatg gaggacgaat gcaacagttc atgctggcct tgttggtggg ggtgatcagc 240  
 tttgttatcg cggcctgatc aagtgtataa tgaataagta aaacgattgc agt 293

<210> 348  
 <211> 70  
 <212> PRT  
 <213> Conus episcopatus

<400> 348  
 Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15  
 Ser Leu Ile Ala Val Pro Leu Asp Gly Asp Gln His Ala Asp Gln Pro  
 20 25 30  
 Ala Glu Arg Leu Gln Gly Asp Ile Leu Ser Glu Lys His Pro Leu Phe  
 35 40 45  
 Met Pro Val Lys Arg Cys Cys Asp Glu Asp Glu Cys Asn Ser Ser Cys  
 50 55 60  
 Trp Pro Cys Cys Trp Gly  
 65 70

<210> 349  
 <211> 16  
 <212> PRT  
 <213> Conus episcopatus

<220>  
 <221> PEPTIDE  
 <222> (1)..(16)  
 <223> Xaa at residue 4 and 6 is Glu or gamma-carboxy Glu; Xaa at residue 13 is Pro or Hyp; Xaa at residue 12 and 16 is Trp or bromo-Trp

<400> 349  
 Cys Cys Asp Xaa Asp Xaa Cys Asn Ser Ser Cys Xaa Xaa Cys Cys Xaa  
 1 5 10 15

<210> 350  
 <211> 293  
 <212> DNA  
 <213> Conus episcopatus

<400> 350  
 caagagggat cgatagcagt tcatgatgtc taaactggga gtcttggtga ccattctgtct 60  
 acttctgttt tcccttattg ctgttccgct tgatggagat caacatgcag accaacctgc 120  
 agagcgtctg cagggcgaca ttttatctga aaagcatccc ttatttatgc ctgtcaaacg 180  
 gtgttgcatg gaggacgaat gcagcagttc atgctggcct tgttggtggg gatgagcagc 240  
 tttgttatcg cggcctcatc aagtgtctaa tgaataagta aaacgattgc agt 293

<210> 351  
 <211> 70

100

&lt;212&gt; PRT

&lt;213&gt; Conus episcopatus

&lt;400&gt; 351

Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15

Ser Leu Ile Ala Val Pro Leu Asp Gly Asp Gln His Ala Asp Gln Pro  
 20 25 30

Ala Glu Arg Leu Gln Gly Asp Ile Leu Ser Glu Lys His Pro Leu Phe  
 35 40 45

Met Pro Val Lys Arg Cys Cys Asp Glu Asp Glu Cys Ser Ser Ser Cys  
 50 55 60

Trp Pro Cys Cys Trp Gly  
 65 70

&lt;210&gt; 352

&lt;211&gt; 16

&lt;212&gt; PRT

&lt;213&gt; Conus episcopatus

&lt;220&gt;

&lt;221&gt; PEPTIDE

&lt;222&gt; (1)..(16)

&lt;223&gt; Xaa at residue 4 and 6 is Glu or gamma-carboxy Glu; Xaa at residue 13 is Pro or Hyp; Xaa at residue 12 and 16 is Trp or bromo-Trp

&lt;400&gt; 352

Cys Cys Asp Xaa Asp Xaa Cys Ser Ser Ser Cys Xaa Xaa Cys Cys Xaa  
 1 5 10 15

&lt;210&gt; 353

&lt;211&gt; 290

&lt;212&gt; DNA

&lt;213&gt; Conus episcopatus

&lt;400&gt; 353

caagagggat cgatagcagt tcatgatgtc taaactggga gtcttggtga ccatctgtct 60

acttctgttt tcccttactg ctgttccgct tgatggagat caacatgcag accaacctgc 120

agagcgtctg caggcgacaca ttttatctga aaagcatccc ttatttaate ccgtaaacg 180

gtgttgcccg gcggcgccat gtgccatggg atgcaagcct tgttggtgat gagcagcttt 240

gttatcgtgg cctcatcaag tgtctaataa ataagtaaaa cgattgcagt 290

&lt;210&gt; 354

&lt;211&gt; 69

&lt;212&gt; PRT

&lt;213&gt; Conus episcopatus

&lt;400&gt; 354

Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15

Ser Leu Thr Ala Val Pro Leu Asp Gly Asp Gln His Ala Asp Gln Pro  
 20 25 30

Ala Glu Arg Leu Gln Gly Asp Ile Leu Ser Glu Lys His Pro Leu Phe  
 35 40 45

101

Asn Pro Val Lys Arg Cys Cys Pro Ala Ala Ala Cys Ala Met Gly Cys  
 50 55 60

Lys Pro Cys Cys Gly  
 65

<210> 355  
 <211> 15  
 <212> PRT  
 <213> Conus episcopatus

<220>  
 <221> PEPTIDE  
 <222> (1)..(15)  
 <223> Xaa at residue 3 and 13 is Pro or Hyp

<400> 355  
 Cys Cys Xaa Ala Ala Ala Cys Ala Met Gly Cys Lys Xaa Cys Cys  
 1 5 10 15

<210> 356  
 <211> 295  
 <212> DNA  
 <213> Conus aulicus

<400> 356  
 caagagggat cgatagcagt tcatgatgtc taaactggga gtcttggtga coactctgtct 60  
 gcttctgttt tccgttactg ctcttccgcc ggatggagat caacctgcag accgagctgc 120  
 agagcgtagg caggtcgcagc agcatcccgt gtttgatcat gaaagaggggt gttgctcgcc 180  
 accatgccac agtattttgcg ctgctttctg ttgcgggtga tgataacgtg ttgatgaccc 240  
 actttgtcat cacggctgcg tcaagtgtct aatgaataag taaaatgatt gcagt 295

<210> 357  
 <211> 65  
 <212> PRT  
 <213> Conus aulicus

<400> 357  
 Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15

Ser Val Thr Ala Leu Pro Pro Asp Gly Asp Gln Pro Ala Asp Arg Ala  
 20 25 30

Ala Glu Arg Arg Gln Val Glu Gln His Pro Val Phe Asp His Glu Arg  
 35 40 45

Gly Cys Cys Ser Pro Pro Cys His Ser Ile Cys Ala Ala Phe Cys Cys  
 50 55 60

Gly  
 65

<210> 358  
 <211> 16  
 <212> PRT  
 <213> Conus aulicus

<220>  
 <221> PEPTIDE  
 <222> (1)..(16)  
 <223> Xaa at residue 5 and 6 is Pro or Hyp



102

&lt;400&gt; 358

Gly Cys Cys Ser Xaa Xaa Cys His Ser Ile Cys Ala Ala Phe Cys Cys  
 1 5 10 15

&lt;210&gt; 359

&lt;211&gt; 290

&lt;212&gt; DNA

&lt;213&gt; Conus aulicus

&lt;400&gt; 359

caagagggat cgatagcagt tcatgatgtc taaactggga gtcttggtga ccatctgtct 60  
 acttctgttt tcccttactg ctgttccgct tgatggagat caacatgcag accaacctgc 120  
 agagcgtctg cagggcgaca ttttatctga aaagcatccc ttatttaatc ccgtcaaacg 180  
 gtgttgccga ccggtggcat gtgccatggg atgcaagcct tgttggtgat gaggagcttt 240  
 gttatcgtgg cctcatcaag tgtctaataa ataagtaaaa tgattgcagt 290

&lt;210&gt; 360

&lt;211&gt; 69

&lt;212&gt; PRT

&lt;213&gt; Conus aulicus

&lt;400&gt; 360

Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15

Ser Leu Thr Ala Val Pro Leu Asp Gly Asp Gln His Ala Asp Gln Pro  
 20 25 30

Ala Glu Arg Leu Gln Gly Asp Ile Leu Ser Glu Lys His Pro Leu Phe  
 35 40 45

Asn Pro Val Lys Arg Cys Cys Arg Pro Val Ala Cys Ala Met Gly Cys  
 50 55 60

Lys Pro Cys Cys Gly  
 65

&lt;210&gt; 361

&lt;211&gt; 15

&lt;212&gt; PRT

&lt;213&gt; Conus aulicus

&lt;220&gt;

&lt;221&gt; PEPTIDE

&lt;222&gt; (1)..(15)

&lt;223&gt; Xaa at residue 4 and 13 is Pro or Hyp

&lt;400&gt; 361

Cys Cys Arg Xaa Val Ala Cys Ala Met Gly Cys Lys Xaa Cys Cys  
 1 5 10 15

&lt;210&gt; 362

&lt;211&gt; 290

&lt;212&gt; DNA

&lt;213&gt; Conus aulicus

&lt;400&gt; 362

caagagggat cgatagcagt tcatgatgtc taaactggga gtcttggtga tcatctgtct 60  
 acttctgtct ccccttactg ctgttccgct ggatggagat caacctgcag accgacctgc 120

agagcgtatg caggacgaca ttcatctga acatcaaccc atgtttgatg ccatcagaca 180  
 gtgttgcccg gcggtggcat gcgccatggg atgcgagcct tgttgtggat gaccagcttt 240  
 gttatcgcg cctcatcaag tgtctaata gaataaaaa tgattgcagt 290

<210> 363  
 <211> 69  
 <212> PRT  
 <213> Conus aulicus

<400> 363  
 Met Met Ser Lys Leu Gly Val Leu Leu Ile Ile Cys Leu Leu Leu Ser  
 1 5 10 15  
 Pro Leu Thr Ala Val Pro Leu Asp Gly Asp Gln Pro Ala Asp Arg Pro  
 20 25 30  
 Ala Glu Arg Met Gln Asp Asp Ile Ser Ser Glu His Gln Pro Met Phe  
 35 40 45  
 Asp Ala Ile Arg Gln Cys Cys Pro Ala Val Ala Cys Ala Met Gly Cys  
 50 55 60  
 Glu Pro Cys Cys Gly  
 65

<210> 364  
 <211> 16  
 <212> PRT  
 <213> Conus aulicus

<220>  
 <221> PEPTIDE  
 <222> (1)..(16)  
 <223> Xaa at residue 1 is Gln or pyro-Glu; Xaa at residue 13 is Glu or  
 gamma-carboxy Glu; Xaa at residue 4 and 14 is Pro or Hy

<400> 364  
 Xaa Cys Cys Xaa Ala Val Ala Cys Ala Met Gly Cys Xaa Xaa Cys Cys  
 1 5 10 15

<210> 365  
 <211> 293  
 <212> DNA  
 <213> Conus aureus

<400> 365  
 caagaaggat cgatagcagt tcatgatgtc taaactggga gccttggtga ccatctgtct 60  
 acttctgttt tcccttactg ctgttccgct ggatggagat caacatgcag accaacaatgc 120  
 agagcgtctg catgaccgcc ttccaactga aaatcatccc ttatatgatc ccgtcaaacg 180  
 gtgttgcatg gattcggaat gcgactatc ttgctggcct tgctgtatct ttggataacc 240  
 ttgttatcgc cggcctcatc aagtgtcaaa tgaataagta aaacgattgc agt 293

<210> 366  
 <211> 71  
 <212> PRT  
 <213> Conus aureus

<400> 366

104

Met Met Ser Lys Leu Gly Ala Leu Leu Thr Ile Cys Leu Leu Leu Phe  
1 5 10 15

Ser Leu Thr Ala Val Pro Leu Asp Gly Asp Gln His Ala Asp Gln His  
20 25 30

Ala Glu Arg Leu His Asp Arg Leu Pro Thr Glu Asn His Pro Leu Tyr  
35 40 45

Asp Pro Val Lys Arg Cys Cys Asp Asp Ser Glu Cys Asp Tyr Ser Cys  
50 55 60

Trp Pro Cys Cys Ile Phe Gly  
65 70

<210> 367

<211> 17

<212> PRT

<213> Conus aureus

<220>

<221> PEPTIDE

<222> (1)..(17)

<223> Xaa at residue 6 is Glu or gamma-carboxy Glu; Xaa at residue 13 is Pro or Hyp; Xaa at residue 12 is Trp or bromo-Trp; Xaa at residue 9 is Tyr, 125I-Tyr, mono-iodo-Tyr, di-iodo-Tyr, O-sulpho-Tyr or O-phospho-Ty

<400> 367

Cys Cys Asp Asp Ser Xaa Cys Asp Xaa Ser Cys Xaa Xaa Cys Cys Ile  
1 5 10 15

Phe

<210> 368

<211> 290

<212> DNA

<213> Conus aureus

<400> 368

caagagggat cgaatagcagt tcatgatgtc taaactggga gccttggtga ccatctgtct 60

acttctgttt tccctaactg ctgttcgct ggaaggagat caacatgcag accaacctgc 120

agagcgtctg caggaccgca ttccaactga aaatcatccc ttatttgatc cgaacaaacg 180

gtgttgcaat gattgggaat gcgacgattc atgtggcct tgctgttatg gataaccttt 240

gttatcgcg cctcatcaag tgtcaaatga ataagtaaaa cgattgcagt 290

<210> 369

<211> 70

<212> PRT

<213> Conus aureus

<400> 369

Met Met Ser Lys Leu Gly Ala Leu Leu Thr Ile Cys Leu Leu Leu Phe  
1 5 10 15

Ser Leu Thr Ala Val Pro Leu Asp Gly Asp Gln His Ala Asp Gln Pro  
20 25 30

Ala Glu Arg Leu Gln Asp Arg Ile Pro Thr Glu Asn His Pro Leu Phe  
35 40 45

105

Asp Pro Asn Lys Arg Cys Cys Asn Asp Trp Glu Cys Asp Asp Ser Cys  
 50 55 60

Trp Pro Cys Cys Tyr Gly  
 65 70

<210> 370  
 <211> 16  
 <212> PRT  
 <213> Conus aureus

<220>  
 <221> PEPTIDE  
 <222> (1)..(16)  
 <223> Xaa at residue 6 is Glu or gamma-carboxy Glu; Xaa at residue 13 is Pro or Hyp; Xaa at residue 5 and 12 is Trp or bromo-Trp; Xaa at residue 16 is Tyr, 125I-Tyr, mono-iodo-Tyr, di-iodo-Tyr, O-sulpho-Tyr or O-phospho-Ty

<400> 370  
 Cys Cys Asn Asp Xaa Xaa Cys Asp Asp Ser Cys Xaa Xaa Cys Cys Xaa  
 1 5 10 15

<210> 371  
 <211> 310  
 <212> DNA  
 <213> Conus consors

<400> 371  
 caagagggat cgatagcagt tcatgatgtc taaactggga gtcttggtga ccatctgttt 60  
 gcttctgttt ccccttactg ctcttccaat ggatggagat caatctgtag accgacctgc 120  
 agagcgtatg caggacgaca tttcatctga gctgcatccc ttgttcaatc agaaaagaat 180  
 gtgttgcggc gaaggtgcmc catgccccag ctatttcaga aacagtcaga tttgtcattg 240  
 ttgttaaatg acaacgtgtc gatgaccaac ttogttatca cgactaatga ataagtaaaa 300  
 tgattgcagt 310

<210> 372  
 <211> 74  
 <212> PRT  
 <213> Conus consors

<400> 372  
 Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15

Pro Leu Thr Ala Leu Pro Met Asp Gly Asp Gln Ser Val Asp Arg Pro  
 20 25 30

Ala Glu Arg Met Gln Asp Asp Ile Ser Ser Glu Leu His Pro Leu Phe  
 35 40 45

Asn Gln Lys Arg Met Cys Cys Gly Glu Gly Ala Pro Cys Pro Ser Tyr  
 50 55 60

Phe Arg Asn Ser Gln Ile Cys His Cys Cys  
 65 70

<210> 373  
 <211> 22  
 <212> PRT

106

&lt;213&gt; Conus consors

&lt;220&gt;

&lt;221&gt; PEPTIDE

&lt;222&gt; (1)..(22)

<223> Xaa at residue 5 is Glu or gamma-carboxy Glu; Xaa at residue 8 and 10 is Pro or Hyp; Xaa at residue 12 is Tyr, 125I-Tyr, mono-iodo-Tyr, di-iodo-Tyr, O-sulpho-Tyr or O-phospho-Ty

&lt;400&gt; 373

Met Cys Cys Gly Xaa Gly Ala Xaa Cys Xaa Ser Xaa Phe Arg Asn Ser  
 1 5 10 15

Gln Ile Cys His Cys Cys  
 20

&lt;210&gt; 374

&lt;211&gt; 315

&lt;212&gt; DNA

&lt;213&gt; Conus consors

&lt;400&gt; 374

taagagggat cgatagcagt tcatgatgtc taaactggga gtcttggtga ccatctgtct 60  
 gcttctgttt ccccttattg ctcttccaat ggatggagat caacctgcag accgacctgc 120  
 agagcgtatg caggacgaca ttcatctca gcagcatccc ttgtttgata agagaggccg 180  
 ctgttgcgat gtgccgaacg catgctccgg cagatggtgc agagatcacg cacaatgttg 240  
 cggatgacga taacgtgttg atgaccaact ttgtgatcac ggctacatca agtgaataag 300  
 taaaacgatt gcagt 315

&lt;210&gt; 375

&lt;211&gt; 74

&lt;212&gt; PRT

&lt;213&gt; Conus consors

&lt;400&gt; 375

Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15

Pro Leu Ile Ala Leu Pro Met Asp Gly Asp Gln Pro Ala Asp Arg Pro  
 20 25 30

Ala Glu Arg Met Gln Asp Asp Ile Ser Ser Gln Gln His Pro Leu Phe  
 35 40 45

Asp Lys Arg Gly Arg Cys Cys Asp Val Pro Asn Ala Cys Ser Gly Arg  
 50 55 60

Trp Cys Arg Asp His Ala Gln Cys Cys Gly  
 65 70

&lt;210&gt; 376

&lt;211&gt; 22

&lt;212&gt; PRT

&lt;213&gt; Conus consors

&lt;220&gt;

&lt;221&gt; PEPTIDE

&lt;222&gt; (1)..(22)

<223> Xaa at residue 7 is Pro or Hyp; Xaa at residue 14 is Trp or bromo-Tr

107

&lt;400&gt; 376

Gly Arg Cys Cys Asp Val Xaa Asn Ala Cys Ser Gly Arg Xaa Cys Arg  
 1 5 10 15

Asp His Ala Gln Cys Cys  
 20

&lt;210&gt; 377

&lt;211&gt; 322

&lt;212&gt; DNA

&lt;213&gt; Conus consors

&lt;400&gt; 377

caagagggat cgatagcagt tcatgatgtc taaactggga gtcttggtga ctgtctgttt 60  
 gcttctgttt ccccttactg ctcttccgat ggatggagat caacctgcag accaacctgc 120  
 agagcgtatg caggacgaca ttcatctga gcagcatccc ttgtttgata agagacaaag 180  
 gtgttgcaact gggaagaagg ggatcatgtc cggtaaagca tgcaaaagtc tcaaatgttg 240  
 ctctggacga taacgtgttg atgaccaact ttgttatcac ggctacgtca agtgtctagt 300  
 gaataagtaa aacgattgca gt 322

&lt;210&gt; 378

&lt;211&gt; 76

&lt;212&gt; PRT

&lt;213&gt; Conus consors

&lt;400&gt; 378

Met Met Ser Lys Leu Gly Val Leu Leu Thr Val Cys Leu Leu Leu Phe  
 1 5 10 15

Pro Leu Thr Ala Leu Pro Met Asp Gly Asp Gln Pro Ala Asp Gln Pro  
 20 25 30

Ala Glu Arg Met Gln Asp Asp Ile Ser Ser Glu Gln His Pro Leu Phe  
 35 40 45

Asp Lys Arg Gln Arg Cys Cys Thr Gly Lys Lys Gly Ser Cys Ser Gly  
 50 55 60

Lys Ala Cys Lys Ser Leu Lys Cys Cys Ser Gly Arg  
 65 70 75

&lt;210&gt; 379

&lt;211&gt; 23

&lt;212&gt; PRT

&lt;213&gt; Conus consors

&lt;220&gt;

&lt;221&gt; PEPTIDE

&lt;222&gt; (1)..(23)

&lt;223&gt; Xaa at residue 1 is Gln or pyro-Glu

&lt;400&gt; 379

Xaa Arg Cys Cys Thr Gly Lys Lys Gly Ser Cys Ser Gly Lys Ala Cys  
 1 5 10 15

Lys Ser Leu Lys Cys Cys Ser  
 20

&lt;210&gt; 380

108

<211> 284  
 <212> DNA  
 <213> Conus emaciatus

<400> 380  
 caagagggat cgatagcagt tcatgatgtc taaactggga gtcttgctga ccatctgtct 60  
 gcttctgttt cccettactg ttcttcgat ggatggagat caacctgcag acctacctgc 120  
 attgctgctg cagttctttg cacctgaaca tagtccccgg ttgacccccg tcaaacggtg 180  
 ctgctcgcgg gattgcagtg ttgcatccc ttgttgcccg tatggatcac cttgattatt 240  
 gcggccacgt caagtgtcta atgaataagt aaaatgattg cagt 284

<210> 381  
 <211> 70  
 <212> PRT  
 <213> Conus emaciatus

<400> 381  
 Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15  
 Pro Leu Thr Val Leu Pro Met Asp Gly Asp Gln Pro Ala Asp Leu Pro  
 20 25 30  
 Ala Leu Arg Ala Gln Phe Phe Ala Pro Glu His Ser Pro Arg Phe Asp  
 35 40 45  
 Pro Val Lys Arg Cys Cys Ser Arg Asp Cys Ser Val Cys Ile Pro Cys  
 50 55 60  
 Cys Pro Tyr Gly Ser Pro  
 65 70

<210> 382  
 <211> 18  
 <212> PRT  
 <213> Conus emaciatus

<220>  
 <221> PEPTIDE  
 <222> (1)..(18)  
 <223> Xaa at residue 11, 14 and 18 is Pro or Hyp; Xaa at residue 15 is  
 Tyr, 125I-Tyr, mono-iodo-Tyr, di-iodo-Tyr, O-sulpho-Tyr or O-phos  
 pho-Ty

<400> 382  
 Cys Cys Ser Arg Asp Cys Ser Val Cys Ile Xaa Cys Cys Xaa Xaa Gly  
 1 5 10 15

Ser Xaa

<210> 383  
 <211> 13  
 <212> PRT  
 <213> Conus aurisiacus

<400> 383  
 Cys Cys Lys Val Gln Cys Glu Ser Cys Thr Pro Cys Cys  
 1 5 10

<210> 384  
 <211> 15

109

<212> PRT  
 <213> Conus atlanticus

<400> 384  
 Cys Cys Glu Leu Pro Cys Gly Pro Gly Phe Cys Val Pro Cys Cys  
 1 5 10 15

<210> 385  
 <211> 14  
 <212> PRT  
 <213> Conus arentus

<400> 385  
 Cys Cys Glu Arg Pro Cys Asn Ile Gly Cys Val Pro Cys Cys  
 1 5 10

<210> 386  
 <211> 16  
 <212> PRT  
 <213> Conus bandus

<400> 386  
 Cys Cys Asn Trp Pro Cys Ser Met Gly Cys Ile Pro Cys Cys Tyr Tyr  
 1 5 10 15

<210> 387  
 <211> 15  
 <212> PRT  
 <213> Conus betulinus

<400> 387  
 Cys Cys Glu Leu Pro Cys His Gly Cys Val Pro Cys Cys Trp Pro  
 1 5 10 15

<210> 388  
 <211> 16  
 <212> PRT  
 <213> Conus betulinus

<400> 388  
 Cys Cys Gly Leu Pro Cys Asn Gly Cys Val Pro Cys Cys Trp Pro Ser  
 1 5 10 15

<210> 389  
 <211> 18  
 <212> PRT  
 <213> Conus betulinus

<400> 389  
 Cys Cys Ser Arg Asn Cys Ala Val Cys Ile Pro Cys Cys Pro Asn Trp  
 1 5 10 15

Pro Ala

<210> 390  
 <211> 14  
 <212> PRT  
 <213> Conus betulinus

<400> 390  
 Cys Cys Lys Gln Ser Cys Thr Thr Cys Met Pro Cys Cys Trp  
 1 5 10

<210> 391  
 <211> 14



110

<212> PRT  
<213> Conus betulinus

<220>  
<221> PEPTIDE  
<222> (1)..(14)  
<223> Xaa is Glu or gamma-carboxy Glu

<400> 391  
Ala Cys Cys Xaa Gln Ser Cys Thr Thr Cys Met Pro Cys Cys  
1 5 10

<210> 392  
<211> 14  
<212> PRT  
<213> Conus betulinus

<400> 392  
Cys Cys Glu Gln Ser Cys Thr Thr Cys Met Pro Cys Cys Trp  
1 5 10

<210> 393  
<211> 18  
<212> PRT  
<213> Conus characteristicus

<400> 393  
Arg Cys Cys Arg Tyr Pro Cys Pro Asp Ser Cys His Gly Ser Cys Cys  
1 5 10 15

Tyr Lys

<210> 394  
<211> 15  
<212> PRT  
<213> Conus characteristicus

<400> 394  
Cys Cys Pro Pro Val Ala Cys Asn Met Gly Cys Lys Pro Cys Cys  
1 5 10 15

<210> 395  
<211> 17  
<212> PRT  
<213> Conus characteristicus

<400> 395  
Cys Cys Asp Asp Ser Glu Cys Asp Tyr Ser Cys Trp Pro Cys Cys Met  
1 5 10 15

Phe

<210> 396  
<211> 14  
<212> PRT  
<213> Conus characteristicus

<400> 396  
Cys Cys Arg Arg Cys Tyr Met Gly Cys Ile Pro Cys Cys Phe  
1 5 10

<210> 397  
<211> 16  
<212> PRT  
<213> Conus textile

111

<400> 397  
 Cys Cys Pro Pro Val Ala Cys Asn Met Gly Cys Lys Pro Cys Cys Gly  
 1 5 10 15

<210> 398  
 <211> 19  
 <212> PRT  
 <213> Conus marmoreus

<220>  
 <221> PEPTIDE  
 <222> (1)..(19)  
 <223> Xaa is Hyp

<400> 398  
 Ser Lys Gln Cys Cys His Leu Ala Ala Cys Arg Phe Gly Cys Thr Xaa  
 1 5 10 15

Cys Cys Asn

<210> 399  
 <211> 15  
 <212> PRT  
 <213> Conus capitaneus

<400> 399  
 Ser Cys Cys Arg Asp Cys Gly Glu Asp Cys Val Gly Cys Cys Arg  
 1 5 10 15

<210> 400  
 <211> 16  
 <212> PRT  
 <213> Conus coronatus

<400> 400  
 Cys Cys Asp Trp Pro Cys Ile Pro Gly Cys Thr Pro Cys Cys Leu Pro  
 1 5 10 15

<210> 401  
 <211> 18  
 <212> PRT  
 <213> Conus dalli

<400> 401  
 Cys Cys Asp Asp Ser Glu Cys Asp Tyr Ser Cys Trp Pro Cys Cys Ile  
 1 5 10 15

Leu Ser

<210> 402  
 <211> 17  
 <212> PRT  
 <213> Conus dalli

<400> 402  
 Glx Gln Cys Cys Pro Pro Val Ala Cys Asn Met Gly Cys Glu Pro Cys  
 1 5 10 15

Cys

<210> 403  
 <211> 16  
 <212> PRT  
 <213> Conus dalli

112

<400> 403  
 Cys Cys Asn Ala Gly Phe Cys Arg Phe Gly Cys Thr Pro Cys Cys Trp  
 1 5 10 15

<210> 404  
 <211> 14  
 <212> PRT  
 <213> Conus distans

<400> 404  
 Glx Cys Cys Val His Pro Cys Pro Cys Thr Pro Cys Cys Arg  
 1 5 10

<210> 405  
 <211> 14  
 <212> PRT  
 <213> Conus figulinus

<400> 405  
 Cys Cys Pro Trp Pro Cys Asn Ile Gly Cys Val Pro Cys Cys  
 1 5 10

<210> 406  
 <211> 14  
 <212> PRT  
 <213> Conus figulinus

<400> 406  
 Cys Cys Ser Lys Asn Cys Ala Val Cys Ile Pro Cys Cys Pro  
 1 5 10

<210> 407  
 <211> 15  
 <212> PRT  
 <213> Conus figulinus

<400> 407  
 Cys Cys Arg Trp Pro Cys Pro Ala Arg Cys Gly Ser Cys Cys Leu  
 1 5 10 15

<210> 408  
 <211> 16  
 <212> PRT  
 <213> Conus figulinus

<400> 408  
 Cys Cys Glu Leu Ser Arg Cys Leu Gly Cys Val Pro Cys Cys Thr Ser  
 1 5 10 15

<210> 409  
 <211> 16  
 <212> PRT  
 <213> Conus figulinus

<400> 409  
 Cys Cys Glu Leu Ser Lys Cys His Gly Cys Val Pro Cys Cys Ile Pro  
 1 5 10 15

<210> 410  
 <211> 16  
 <212> PRT  
 <213> Conus generalis

<400> 410

113

Glx Cys Cys Thr Phe Cys Asn Phe Gly Cys Gln Pro Cys Cys Val Pro  
 1 5 10 15

<210> 411  
 <211> 16  
 <212> PRT  
 <213> Conus generalis

<400> 411  
 Glx Cys Cys Thr Phe Cys Asn Phe Gly Cys Gln Pro Cys Cys Leu Thr  
 1 5 10 15

<210> 412  
 <211> 16  
 <212> PRT  
 <213> Conus generalis

<400> 412

Glx Cys Cys Thr Phe Cys Asn Phe Gly Cys Gln Pro Cys Cys Val Pro  
 1 5 10 15

<210> 413  
 <211> 17  
 <212> PRT  
 <213> Conus gloriamaris

<400> 413  
 Cys Cys Asp Asp Ser Glu Cys Asp Tyr Ser Cys Trp Pro Cys Cys Met  
 1 5 10 15

Phe

<210> 414  
 <211> 17  
 <212> PRT  
 <213> Conus gloriamaris

<400> 414  
 Gly Cys Cys His Leu Leu Ala Cys Arg Phe Gly Cys Ser Pro Cys Cys  
 1 5 10 15  
 Trp

<210> 415  
 <211> 16  
 <212> PRT  
 <213> Conus gloriamaris

<400> 415  
 Cys Cys Ser Trp Asp Val Cys Asp His Pro Ser Cys Thr Cys Cys Gly  
 1 5 10 15

<210> 416  
 <211> 13  
 <212> PRT  
 <213> Conus laterculatus

<400> 416  
 Cys Cys Asp Trp Pro Cys Ser Gly Cys Ile Pro Cys Cys  
 1 5 10

<210> 417  
 <211> 19  
 <212> PRT  
 <213> Conus leopardus

114

&lt;400&gt; 417

Glx Ile Asn Cys Cys Pro Trp Pro Cys Pro Ser Thr Cys Arg His Gln  
1 5 10 15

Cys Cys His

&lt;210&gt; 418

&lt;211&gt; 19

&lt;212&gt; PRT

&lt;213&gt; Conus lividus

&lt;400&gt; 418

Glx Ile Asn Cys Cys Pro Trp Pro Cys Pro Asp Ser Cys His Tyr Gln  
1 5 10 15

Cys Cys His

&lt;210&gt; 419

&lt;211&gt; 14

&lt;212&gt; PRT

&lt;213&gt; Conus marmoreus

&lt;400&gt; 419

Cys Cys Arg Leu Ser Cys Gly Leu Gly Cys His Pro Cys Cys  
1 5 10

&lt;210&gt; 420

&lt;211&gt; 17

&lt;212&gt; PRT

&lt;213&gt; Conus marmoreus

&lt;400&gt; 420

Glu Cys Cys Gly Ser Phe Ala Cys Arg Phe Gly Cys Val Pro Cys Cys  
1 5 10 15

Val

&lt;210&gt; 421

&lt;211&gt; 19

&lt;212&gt; PRT

&lt;213&gt; Conus marmoreus

&lt;400&gt; 421

Ser Lys Gln Cys Cys His Leu Pro Ala Cys Arg Phe Gly Cys Thr Pro  
1 5 10 15

Cys Cys Trp

&lt;210&gt; 422

&lt;211&gt; 17

&lt;212&gt; PRT

&lt;213&gt; Conus marmoreus

&lt;400&gt; 422

Met Gly Cys Cys Pro Phe Pro Cys Lys Thr Ser Cys Thr Thr Leu Cys  
1 5 10 15

Cys

&lt;210&gt; 423

&lt;211&gt; 14

&lt;212&gt; PRT

&lt;213&gt; Conus musicus

&lt;400&gt; 423

115

Ala Cys Cys Glu Gln Ser Cys Thr Thr Cys Phe Pro Cys Cys  
 1 5 10

<210> 424  
 <211> 15  
 <212> PRT  
 <213> Conus nobilis

<400> 424  
 Cys Cys Glu Leu Pro Cys Gly Pro Gly Phe Cys Val Pro Cys Cys  
 1 5 10 15

<210> 425  
 <211> 14  
 <212> PRT  
 <213> Conus pulicarius

<400> 425  
 Cys Cys Asn Ser Cys Tyr Met Gly Cys Ile Pro Cys Cys Phe  
 1 5 10

<210> 426  
 <211> 17  
 <212> PRT  
 <213> Conus quercinus

<400> 426  
 Glx Arg Cys Cys Gln Trp Pro Cys Pro Gly Ser Cys Arg Cys Cys Arg  
 1 5 10 15

Thr

<210> 427  
 <211> 18  
 <212> PRT  
 <213> Conus quercinus

<400> 427  
 Glx Arg Cys Cys Arg Trp Pro Cys Pro Gly Ser Cys Arg Cys Cys Arg  
 1 5 10 15

Tyr Arg

<210> 428  
 <211> 18  
 <212> PRT  
 <213> Conus quercinus

<400> 428  
 Arg Cys Cys Arg Tyr Pro Cys Pro Asp Ser Cys His Gly Ser Cys Cys  
 1 5 10 15

Tyr Lys

<210> 429  
 <211> 15  
 <212> PRT  
 <213> Conus quercinus

<220>  
 <221> PEPTIDE  
 <222> (1)..(15)  
 <223> Xaa is Hyp

<400> 429

116

Cys Cys Ser Gln Asp Cys Leu Val Cys Ile Xaa Cys Cys Pro Asn  
 1 5 10 15

<210> 430  
 <211> 15  
 <212> PRT  
 <213> Conus quercinus

<220>  
 <221> PEPTIDE  
 <222> (1)..(15)  
 <223> Xaa is Hyp

<400> 430  
 Cys Cys Ser Arg His Cys Trp Val Cys Ile Xaa Cys Cys Pro Asn  
 1 5 10 15

<210> 431  
 <211> 16  
 <212> PRT  
 <213> Conus rattus

<400> 431  
 Glx Thr Cys Cys Ser Asn Cys Gly Glu Asp Cys Asp Gly Cys Cys Gln  
 1 5 10 15

<210> 432  
 <211> 20  
 <212> PRT  
 <213> Conus striatus

<400> 432  
 Glx Asn Cys Cys Asn Gly Gly Cys Ser Ser Lys Trp Cys Arg Asp His  
 1 5 10 15

Ala Arg Cys Cys  
 20

<210> 433  
 <211> 12  
 <212> PRT  
 <213> Conus textile

<220>  
 <221> PEPTIDE  
 <222> (1)..(12)  
 <223> Xaa is Hyp

<400> 433  
 Cys Cys Arg Thr Cys Phe Gly Cys Thr Xaa Cys Cys  
 1 5 10

<210> 434  
 <211> 14  
 <212> PRT  
 <213> Conus tessulatus

<400> 434  
 Cys Cys His Lys Cys Tyr Met Gly Cys Ile Pro Cys Cys Ile  
 1 5 10

<210> 435  
 <211> 18  
 <212> PRT  
 <213> Conus tessulatus

117

&lt;400&gt; 435

Lys Cys Cys Arg Pro Pro Cys Ala Met Ser Cys Gly Met Ala Arg Cys  
1 5 10 15

Cys Tyr

&lt;210&gt; 436

&lt;211&gt; 23

&lt;212&gt; PRT

&lt;213&gt; Conus betulinus

&lt;400&gt; 436

Arg Cys Cys Arg Trp Pro Cys Pro Ser Ile Cys Gly Met Ala Arg Cys  
1 5 10 15

Cys Phe Val Met Ile Thr Cys  
20

&lt;210&gt; 437

&lt;211&gt; 23

&lt;212&gt; PRT

&lt;213&gt; Conus betulinus

&lt;400&gt; 437

Arg Cys Cys Arg Trp Pro Cys Pro Ser Arg Cys Gly Met Ala Arg Cys  
1 5 10 15

Cys Phe Val Met Ile Thr Cys  
20

&lt;210&gt; 438

&lt;211&gt; 15

&lt;212&gt; PRT

&lt;213&gt; Conus textile

&lt;400&gt; 438

Phe Cys Cys Asp Ser Asn Trp Cys His Asp Cys Glu Cys Cys Tyr  
1 5 10 15

&lt;210&gt; 439

&lt;211&gt; 16

&lt;212&gt; PRT

&lt;213&gt; Conus marmoreus

&lt;400&gt; 439

Cys Cys His Trp Asn Trp Cys Asp His Leu Cys Ser Cys Cys Gly Ser  
1 5 10 15

&lt;210&gt; 440

&lt;211&gt; 16

&lt;212&gt; PRT

&lt;213&gt; Conus marmoreus

&lt;220&gt;

&lt;221&gt; PEPTIDE

&lt;222&gt; (1)..(16)

&lt;223&gt; Xaa is Hyp

&lt;400&gt; 440

Asp Cys Cys Xaa Leu Pro Ala Cys Pro Phe Gly Cys Asn Xaa Cys Cys  
1 5 10 15

&lt;210&gt; 441

&lt;211&gt; 16



118

<212> PRT  
<213> Conus marmoreus

<220>  
<221> PEPTIDE  
<222> (1)..(16)  
<223> Xaa is Hyp

<400> 441  
Cys Cys Ala Pro Ser Ala Cys Arg Leu Gly Cys Arg Xaa Cys Cys Arg  
1 5 10 15

<210> 442  
<211> 16  
<212> PRT  
<213> Conus marmoreus

<220>  
<221> PEPTIDE  
<222> (1)..(16)  
<223> Xaa is Hyp

<400> 442  
Cys Cys Ala Xaa Ser Ala Cys Arg Leu Gly Cys Arg Xaa Cys Cys Arg  
1 5 10 15

<210> 443  
<211> 16  
<212> PRT  
<213> Conus marmoreus

<400> 443  
Cys Cys Ala Pro Ser Ala Cys Arg Leu Gly Cys Arg Pro Cys Cys Arg  
1 5 10 15

<210> 444  
<211> 17  
<212> PRT  
<213> Conus marmoreus

<220>  
<221> PEPTIDE  
<222> (1)..(17)  
<223> Xaa is Hyp

<400> 444  
Gly Cys Cys Gly Ser Phe Ala Cys Arg Phe Gly Cys Val Xaa Cys Cys  
1 5 10 15

Val

<210> 445  
<211> 15  
<212> PRT  
<213> Conus textile

<400> 445  
Cys Cys Ser Trp Asp Val Cys Asp His Pro Ser Cys Thr Cys Cys  
1 5 10 15

<210> 446  
<211> 16  
<212> PRT  
<213> Conus textile

119

&lt;400&gt; 446

Arg Cys Cys Lys Phe Pro Cys Pro Asp Ser Cys Arg Tyr Leu Cys Cys  
1 5 10 15

&lt;210&gt; 447

&lt;211&gt; 17

&lt;212&gt; PRT

&lt;213&gt; Conus aureus

&lt;400&gt; 447

Cys Cys Asp Asp Ser Glu Cys Asp Tyr Ser Cys Trp Pro Cys Cys Ile  
1 5 10 15

Phe

&lt;210&gt; 448

&lt;211&gt; 16

&lt;212&gt; PRT

&lt;213&gt; Conus aureus

&lt;400&gt; 448

Cys Cys Asn Asp Trp Glu Cys Asp Asp Ser Cys Trp Pro Cys Cys Tyr  
1 5 10 15

&lt;210&gt; 449

&lt;211&gt; 16

&lt;212&gt; PRT

&lt;213&gt; Conus ammiralis

&lt;400&gt; 449

Arg Cys Cys Arg Phe Pro Cys Pro Asp Thr Cys Arg His Leu Cys Cys  
1 5 10 15

&lt;210&gt; 450

&lt;211&gt; 12

&lt;212&gt; PRT

&lt;213&gt; Conus ammiralis

&lt;400&gt; 450

Cys Cys Met Thr Cys Phe Gly Cys Thr Pro Cys Cys  
1 5 10

&lt;210&gt; 451

&lt;211&gt; 18

&lt;212&gt; PRT

&lt;213&gt; Conus ammiralis

&lt;400&gt; 451

Cys Cys Asp Asp Ser Glu Cys Asp Tyr Ser Cys Trp Pro Cys Cys Ile  
1 5 10 15

Phe Ser

&lt;210&gt; 452

&lt;211&gt; 13

&lt;212&gt; PRT

&lt;213&gt; Conus ammiralis

&lt;400&gt; 452

Cys Cys Arg Leu Leu Cys Leu Ser Cys Asn Pro Cys Cys  
1 5 10

&lt;210&gt; 453

&lt;211&gt; 16

&lt;212&gt; PRT

120

<213> *Conus ammiralis*

&lt;400&gt; 453

Cys	Cys	Asp	Asp	Ser	Glu	Cys	Gly	Tyr	Ser	Cys	Trp	Pro	Cys	Cys	Tyr
1				5					10					15	

&lt;210&gt; 454

&lt;211&gt; 16

&lt;212&gt; PRT

<213> *Conus aulicus*

&lt;400&gt; 454

Gly	Cys	Cys	Ser	Pro	Pro	Cys	His	Ser	Ile	Cys	Ala	Ala	Phe	Cys	Cys
1				5					10					15	

&lt;210&gt; 455

&lt;211&gt; 15

&lt;212&gt; PRT

<213> *Conus aulicus*

&lt;400&gt; 455

Cys	Cys	Arg	Pro	Val	Ala	Cys	Ala	Met	Gly	Cys	Lys	Pro	Cys	Cys
1				5					10					15

&lt;210&gt; 456

&lt;211&gt; 16

&lt;212&gt; PRT

<213> *Conus aulicus*

&lt;400&gt; 456

Glx	Cys	Cys	Pro	Ala	Val	Ala	Cys	Ala	Met	Gly	Cys	Glu	Pro	Cys	Cys
1				5					10					15	

&lt;210&gt; 457

&lt;211&gt; 18

&lt;212&gt; PRT

<213> *Conus emaciatus*

&lt;400&gt; 457

Cys	Cys	Ser	Arg	Asp	Cys	Ser	Val	Cys	Ile	Pro	Cys	Cys	Pro	Tyr	Gly
1				5					10					15	

Ser Pro

&lt;210&gt; 458

&lt;211&gt; 16

&lt;212&gt; PRT

<213> *Conus episcopatus*

&lt;400&gt; 458

Cys	Cys	Asp	Glu	Asp	Glu	Cys	Asn	Ser	Ser	Cys	Trp	Pro	Cys	Cys	Trp
1				5					10					15	

&lt;210&gt; 459

&lt;211&gt; 16

&lt;212&gt; PRT

<213> *Conus episcopatus*

&lt;400&gt; 459

Cys	Cys	Asp	Glu	Asp	Glu	Cys	Ser	Ser	Ser	Cys	Trp	Pro	Cys	Cys	Trp
1				5					10					15	

&lt;210&gt; 460

&lt;211&gt; 15

&lt;212&gt; PRT

121

&lt;213&gt; Conus episcopatus

&lt;400&gt; 460

Cys	Cys	Pro	Ala	Ala	Ala	Cys	Ala	Met	Gly	Cys	Lys	Pro	Cys	Cys
1				5					10					15

&lt;210&gt; 461

&lt;211&gt; 16

&lt;212&gt; PRT

&lt;213&gt; Conus omaria

&lt;400&gt; 461

Cys	Cys	Asp	Glu	Glu	Glu	Cys	Ser	Ser	Ala	Cys	Trp	Pro	Cys	Cys	Trp
1				5					10						15

&lt;210&gt; 462

&lt;211&gt; 16

&lt;212&gt; PRT

&lt;213&gt; Conus omaria

&lt;400&gt; 462

Cys	Cys	His	Leu	Leu	Ala	Cys	Arg	Phe	Gly	Cys	Ser	Pro	Cys	Cys	Trp
1				5					10						15

&lt;210&gt; 463

&lt;211&gt; 12

&lt;212&gt; PRT

&lt;213&gt; Conus spurius

&lt;400&gt; 463

Cys	Cys	Pro	Arg	Cys	Ser	Glu	Cys	Asn	Pro	Cys	Cys
1				5					10		

&lt;210&gt; 464

&lt;211&gt; 16

&lt;212&gt; PRT

&lt;213&gt; Conus pennaceus

&lt;400&gt; 464

Arg	Cys	Cys	Lys	Phe	Pro	Cys	Pro	Asp	Ser	Cys	Lys	Tyr	Leu	Cys	Cys
1				5					10						15

&lt;210&gt; 465

&lt;211&gt; 19

&lt;212&gt; PRT

&lt;213&gt; Conus flavidus

&lt;400&gt; 465

Arg	Cys	Cys	Arg	Trp	Pro	Cys	Pro	Ser	Ile	Cys	Gly	Met	Ala	Arg	Cys
1				5					10						15

Cys Ser Ser

&lt;210&gt; 466

&lt;211&gt; 14

&lt;212&gt; PRT

&lt;213&gt; Conus pulicarius

&lt;400&gt; 466

Cys	Cys	Lys	Leu	Leu	Cys	Gly	Cys	Thr	Pro	Cys	Cys	His	Ile
1				5					10				

&lt;210&gt; 467

&lt;211&gt; 15

&lt;212&gt; PRT

122

&lt;213&gt; Conus ebraceus

&lt;400&gt; 467

Cys Cys Glu Gln Pro Cys Tyr Met Gly Cys Ile Pro Cys Cys Phe  
 1 5 10 15

&lt;210&gt; 468

&lt;211&gt; 15

&lt;212&gt; PRT

&lt;213&gt; Conus ebraceus

&lt;400&gt; 468

Cys Cys Ala Gln Pro Cys Tyr Met Gly Cys Ile Pro Cys Cys Phe  
 1 5 10 15

&lt;210&gt; 469

&lt;211&gt; 14

&lt;212&gt; PRT

&lt;213&gt; Conus pulicarius

&lt;400&gt; 469

Cys Cys Val Ser Cys Tyr Met Gly Cys Ile Pro Cys Cys Phe  
 1 5 10

&lt;210&gt; 470

&lt;211&gt; 16

&lt;212&gt; PRT

&lt;213&gt; Conus miliaris

&lt;400&gt; 470

Cys Cys Asp Trp Pro Cys Ser Ala Gly Cys Tyr Pro Cys Cys Phe Pro  
 1 5 10 15

&lt;210&gt; 471

&lt;211&gt; 16

&lt;212&gt; PRT

&lt;213&gt; Conus miliaris

&lt;400&gt; 471

Gly Cys Cys Pro Pro Met Cys Thr Pro Cys Phe Pro Cys Cys Phe Arg  
 1 5 10 15

&lt;210&gt; 472

&lt;211&gt; 23

&lt;212&gt; PRT

&lt;213&gt; Conus rattus

&lt;400&gt; 472

Arg Gly Cys Cys Ala Pro Pro Arg Lys Cys Lys Asp Arg Ala Cys Lys  
 1 5 10 15

Pro Ala Arg Cys Cys Gly Pro  
 20

&lt;210&gt; 473

&lt;211&gt; 22

&lt;212&gt; PRT

&lt;213&gt; Conus stercusmuscarum

&lt;400&gt; 473

Glx Arg Cys Cys Asn Gly Arg Arg Gly Cys Ser Ser Arg Trp Cys Arg  
 1 5 10 15

Asp His Ser Arg Cys Cys  
 20

123

<210> 474  
<211> 22  
<212> PRT  
<213> Conus consors

<400> 474  
Gly Arg Cys Cys Asp Val Pro Asn Ala Cys Ser Gly Arg Trp Cys Arg  
1 5 10 15  
Asp His Ala Gln Cys Cys  
20

<210> 475  
<211> 23  
<212> PRT  
<213> Conus consors

<400> 475  
Glx Arg Cys Cys Thr Gly Lys Lys Gly Ser Cys Ser Gly Lys Ala Cys  
1 5 10 15  
Lys Ser Leu Lys Cys Cys Ser  
20

<210> 476  
<211> 22  
<212> PRT  
<213> Conus aurisiacus

<400> 476  
Met Cys Cys Gly Glu Gly Arg Lys Cys Pro Ser Tyr Phe Arg Asn Ser  
1 5 10 15  
Gln Ile Cys His Cys Cys  
20

<210> 477  
<211> 19  
<212> PRT  
<213> Conus aurisiacus

<400> 477  
Cys Cys Arg Trp Pro Cys Pro Arg Gln Ile Asp Gly Glu Tyr Cys Gly  
1 5 10 15  
Cys Cys Leu

<210> 478  
<211> 22  
<212> PRT  
<213> Conus bullatus

<400> 478  
Arg Cys Cys Gly Glu Gly Leu Thr Cys Pro Arg Tyr Trp Lys Asn Ser  
1 5 10 15  
Gln Ile Cys Ala Cys Cys  
20

<210> 479  
<211> 21  
<212> PRT  
<213> Conus characteristicus  
<400> 479

124

Cys Cys Gly Pro Gly Gly Ser Cys Pro Val Tyr Phe Arg Asp Asn Phe  
 1 5 10 15

Ile Cys Gly Cys Cys  
 20

<210> 480  
 <211> 23  
 <212> PRT  
 <213> Conus circumciscus

<400> 480  
 Arg Lys Cys Cys Gly Lys Asp Gly Pro Cys Pro Lys Tyr Phe Lys Asp  
 1 5 10 15

Asn Phe Ile Cys Gly Cys Cys  
 20

<210> 481  
 <211> 20  
 <212> PRT  
 <213> Conus ermineus

<400> 481  
 Cys Cys Ser Trp Pro Cys Pro Arg Tyr Ser Asn Gly Lys Leu Val Cys  
 1 5 10 15

Phe Cys Cys Leu  
 20

<210> 482  
 <211> 21  
 <212> PRT  
 <213> Conus magus

<400> 482  
 Cys Cys Gly Pro Gly Gly Ser Cys Pro Val Tyr Phe Arg Asp Asn Phe  
 1 5 10 15

Ile Cys Gly Cys Cys  
 20

<210> 483  
 <211> 22  
 <212> PRT  
 <213> Conus magus

<400> 483  
 Met Cys Cys Gly Glu Ser Ala Pro Cys Pro Ser Tyr Phe Arg Asn Ser  
 1 5 10 15

Gln Ile Cys His Cys Cys  
 20

<210> 484  
 <211> 22  
 <212> PRT  
 <213> Conus magus

<400> 484  
 Glx Lys Cys Cys Gly Pro Gly Gly Ser Cys Pro Val Tyr Phe Thr Asp  
 1 5 10 15

Asn Phe Ile Cys Gly Cys  
 20

125

<210> 485  
<211> 23  
<212> PRT  
<213> Conus magus

<400> 485  
Glx Lys Cys Cys Gly Pro Gly Gly Ser Cys Pro Val Tyr Phe Arg Asp  
1 5 10 15

Asn Phe Ile Cys Gly Cys Cys  
20

<210> 486  
<211> 23  
<212> PRT  
<213> Conus striatus

<400> 486  
Glx Lys Cys Cys Gly Glu Gly Ser Ser Cys Pro Lys Tyr Phe Lys Asn  
1 5 10 15

Asn Phe Ile Cys Gly Cys Cys  
20

<210> 487  
<211> 22  
<212> PRT  
<213> Conus magus

<400> 487  
Glx Lys Cys Cys Ser Gly Gly Ser Cys Pro Leu Tyr Phe Arg Asp Arg  
1 5 10 15

Leu Ile Cys Pro Cys Cys  
20

<210> 488  
<211> 23  
<212> PRT  
<213> Conus stercusmuscarum

<400> 488  
Glx Lys Cys Cys Gly Pro Gly Ala Ser Cys Pro Arg Tyr Phe Lys Asp  
1 5 10 15

Asn Phe Ile Cys Gly Cys Cys  
20

<210> 489  
<211> 22  
<212> PRT  
<213> Conus consors

<400> 489  
Met Cys Cys Gly Glu Gly Ala Pro Cys Pro Ser Tyr Phe Arg Asn Ser  
1 5 10 15

Gln Ile Cys His Cys Cys  
20

<210> 490  
<211> 23  
<212> PRT  
<213> Conus aurisiacus



126

&lt;400&gt; 490

Glx Lys Cys Cys Thr Gly Lys Lys Gly Ser Cys Ser Gly Lys Ala Cys  
1 5 10 15

Lys Asn Leu Lys Cys Cys Ser  
20

&lt;210&gt; 491

&lt;211&gt; 23

&lt;212&gt; PRT

&lt;213&gt; Conus aurisiacus

&lt;400&gt; 491

Glx Lys Cys Cys Thr Gly Arg Lys Gly Ser Cys Ser Gly Lys Ala Cys  
1 5 10 15

Lys Asn Leu Lys Cys Cys Ser  
20

&lt;210&gt; 492

&lt;211&gt; 23

&lt;212&gt; PRT

&lt;213&gt; Conus bullatus

&lt;400&gt; 492

Val Thr Asp Arg Cys Cys Lys Gly Lys Arg Glu Cys Gly Arg Trp Cys  
1 5 10 15

Arg Asp His Ser Arg Cys Cys  
20

&lt;210&gt; 493

&lt;211&gt; 23

&lt;212&gt; PRT

&lt;213&gt; Conus bullatus

&lt;400&gt; 493

Val Gly Asp Arg Cys Cys Lys Gly Lys Arg Gly Cys Gly Arg Trp Cys  
1 5 10 15

Arg Asp His Ser Arg Cys Cys  
20

&lt;210&gt; 494

&lt;211&gt; 24

&lt;212&gt; PRT

&lt;213&gt; Conus bullatus

&lt;400&gt; 494

Val Gly Glu Arg Cys Cys Lys Asn Gly Lys Arg Gly Cys Gly Arg Trp  
1 5 10 15

Cys Arg Asp His Ser Arg Cys Cys  
20

&lt;210&gt; 495

&lt;211&gt; 26

&lt;212&gt; PRT

&lt;213&gt; Conus bullatus

&lt;400&gt; 495

Ile Val Asp Arg Cys Cys Asn Lys Gly Asn Gly Lys Arg Gly Cys Ser  
1 5 10 15

Arg Trp Cys Arg Asp His Ser Arg Cys Cys

127

20

25

&lt;210&gt; 496

&lt;211&gt; 25

&lt;212&gt; PRT

&lt;213&gt; Conus bullatus

&lt;400&gt; 496

Val	Gly	Cys	Cys	Arg	Pro	Lys	Pro	Asn	Gly	Gln	Met	Met	Cys	Asp	Arg
1				5					10					15	

Trp	Cys	Glu	Lys	Asn	Ser	Arg	Cys	Cys
			20					25

&lt;210&gt; 497

&lt;211&gt; 22

&lt;212&gt; PRT

&lt;213&gt; Conus characteristicus

&lt;400&gt; 497

Arg	Asp	Cys	Cys	Thr	Pro	Pro	Lys	Lys	Cys	Lys	Asp	Arg	Gln	Cys	Lys
1				5					10					15	

Pro	Gln	Arg	Cys	Cys	Ala
			20		

&lt;210&gt; 498

&lt;211&gt; 23

&lt;212&gt; PRT

&lt;213&gt; Conus lynceus

&lt;400&gt; 498

Gly	Arg	Asp	Cys	Cys	Thr	Pro	Pro	Arg	Lys	Cys	Arg	Asp	Arg	Ala	Cys
1				5					10					15	

Lys	Pro	Gln	Arg	Cys	Cys	Gly
				20		

&lt;210&gt; 499

&lt;211&gt; 22

&lt;212&gt; PRT

&lt;213&gt; Conus lynceus

&lt;400&gt; 499

Glx	Arg	Leu	Cys	Cys	Gly	Phe	Pro	Lys	Ser	Cys	Arg	Ser	Arg	Gln	Cys
1				5					10					15	

Lys	Pro	His	Arg	Cys	Cys
				20	

&lt;210&gt; 500

&lt;211&gt; 22

&lt;212&gt; PRT

&lt;213&gt; Conus laterculatus

&lt;400&gt; 500

Arg	Asp	Cys	Cys	Thr	Pro	Pro	Lys	Lys	Cys	Arg	Asp	Arg	Gln	Cys	Lys
1				5					10					15	

Pro	Ala	Arg	Cys	Cys	Gly
				20	

&lt;210&gt; 501

&lt;211&gt; 22

&lt;212&gt; PRT

128

&lt;213&gt; Conus laterculatus

&lt;400&gt; 501

Arg Pro Pro Cys Cys Thr Tyr Asp Gly Ser Cys Leu Lys Glu Ser Cys  
1 5 10 15Met Arg Lys Ala Cys Cys  
20

&lt;210&gt; 502

&lt;211&gt; 22

&lt;212&gt; PRT

&lt;213&gt; Conus laterculatus

&lt;400&gt; 502

Arg Pro Pro Cys Cys Thr Tyr Asp Gly Ser Cys Leu Lys Glu Ser Cys  
1 5 10 15Lys Arg Lys Ala Cys Cys  
20

&lt;210&gt; 503

&lt;211&gt; 22

&lt;212&gt; PRT

&lt;213&gt; Conus geographus

&lt;220&gt;

&lt;221&gt; PEPTIDE

&lt;222&gt; (1)..(22)

&lt;223&gt; Xaa is Hyp

&lt;400&gt; 503

Arg Asp Cys Cys Thr Xaa Xaa Lys Lys Cys Lys Asp Arg Gln Cys Lys  
1 5 10 15Xaa Gln Arg Cys Cys Ala  
20

&lt;210&gt; 504

&lt;211&gt; 22

&lt;212&gt; PRT

&lt;213&gt; Conus geographus

&lt;220&gt;

&lt;221&gt; PEPTIDE

&lt;222&gt; (1)..(22)

&lt;223&gt; Xaa is Hyp

&lt;400&gt; 504

Arg Asp Cys Cys Thr Xaa Xaa Arg Lys Cys Lys Asp Arg Arg Cys Lys  
1 5 10 15Xaa Met Lys Cys Cys Ala  
20

&lt;210&gt; 505

&lt;211&gt; 22

&lt;212&gt; PRT

&lt;213&gt; Conus geographus

&lt;220&gt;

&lt;221&gt; PEPTIDE

&lt;222&gt; (1)..(22)

&lt;223&gt; Xaa is Hyp

129

&lt;400&gt; 505

Arg Asp Cys Cys Thr Xaa Xaa Lys Lys Cys Lys Asp Arg Arg Cys Lys  
1 5 10 15

Xaa Leu Lys Cys Cys Ala  
20

&lt;210&gt; 506

&lt;211&gt; 22

&lt;212&gt; PRT

&lt;213&gt; Conus purpurascens

&lt;220&gt;

&lt;221&gt; PEPTIDE

&lt;222&gt; (1)..(22)

&lt;223&gt; Xaa is Hyp

&lt;400&gt; 506

Glx Arg Leu Cys Cys Gly Phe Xaa Lys Ser Cys Arg Ser Arg Gln Cys  
1 5 10 15

Lys Xaa His Arg Cys Cys  
20

&lt;210&gt; 507

&lt;211&gt; 22

&lt;212&gt; PRT

&lt;213&gt; Conus magus

&lt;400&gt; 507

Arg Asp Cys Cys Thr Pro Pro Lys Lys Cys Lys Asp Arg Gln Cys Lys  
1 5 10 15

Pro Gln Arg Cys Cys Ala  
20

&lt;210&gt; 508

&lt;211&gt; 24

&lt;212&gt; PRT

&lt;213&gt; Conus marmoreus

&lt;400&gt; 508

Arg Gly Gly Cys Cys Thr Pro Pro Arg Lys Cys Lys Asp Arg Ala Cys  
1 5 10 15

Lys Pro Ala Arg Cys Cys Gly Pro  
20

&lt;210&gt; 509

&lt;211&gt; 23

&lt;212&gt; PRT

&lt;213&gt; Conus nobilis

&lt;400&gt; 509

Glx Lys Cys Cys Thr Gly Lys Lys Gly Ser Cys Ser Gly Lys Ala Cys  
1 5 10 15

Lys Asn Leu Lys Cys Cys Ser  
20

&lt;210&gt; 510

&lt;211&gt; 24

&lt;212&gt; PRT

&lt;213&gt; Conus parius

130

&lt;400&gt; 510

Arg Gly Gly Cys Cys Thr Pro Pro Lys Lys Cys Lys Asp Arg Ala Cys  
1 5 10 15

Lys Pro Ala Arg Cys Cys Gly Pro  
20

&lt;210&gt; 511

&lt;211&gt; 23

&lt;212&gt; PRT

&lt;213&gt; Conus parius

&lt;400&gt; 511

Arg Gly Cys Cys Thr Pro Pro Arg Lys Cys Lys Asp Arg Ala Cys Lys  
1 5 10 15

Pro Ala Arg Cys Cys Gly Pro  
20

&lt;210&gt; 512

&lt;211&gt; 24

&lt;212&gt; PRT

&lt;213&gt; Conus radiatus

&lt;220&gt;

&lt;221&gt; PEPTIDE

&lt;222&gt; (1)..(24)

&lt;223&gt; Xaa is Hyp

&lt;400&gt; 512

Leu Xaa Ser Cys Cys Ser Leu Asn Leu Arg Leu Cys Xaa Val Xaa Ala  
1 5 10 15

Cys Lys Arg Asn Xaa Cys Cys Thr  
20

&lt;210&gt; 513

&lt;211&gt; 24

&lt;212&gt; PRT

&lt;213&gt; Conus radiatus

&lt;220&gt;

&lt;221&gt; PEPTIDE

&lt;222&gt; (1)..(24)

&lt;223&gt; Xaa is Hyp

&lt;400&gt; 513

Glx Gln Arg Cys Cys Thr Val Lys Arg Ile Cys Xaa Val Xaa Ala Cys  
1 5 10 15

Arg Ser Lys Xaa Cys Cys Lys Ser  
20

&lt;210&gt; 514

&lt;211&gt; 24

&lt;212&gt; PRT

&lt;213&gt; Conus radiatus

&lt;400&gt; 514

Arg Gly Gly Cys Cys Thr Pro Pro Arg Lys Cys Lys Asp Arg Ala Cys  
1 5 10 15

Lys Pro Ala Arg Cys Cys Gly Pro  
20

131

<210> 515  
 <211> 23  
 <212> PRT  
 <213> Conus stercusmuscarum

<400> 515  
 Glx Lys Cys Cys Thr Gly Lys Lys Gly Ser Cys Ser Gly Lys Ala Cys  
 1 5 10 15

Lys Asn Leu Lys Cys Cys Ser  
 20

<210> 516  
 <211> 21  
 <212> PRT  
 <213> Conus tulipa

<220>  
 <221> PEPTIDE  
 <222> (1)..(21)  
 <223> Xaa is Hyp

<400> 516  
 His Gly Cys Cys Lys Gly Xaa Glu Gly Cys Ser Ser Arg Glu Cys Arg  
 1 5 10 15

Xaa Gln His Cys Cys  
 20

<210> 517  
 <211> 21  
 <212> PRT  
 <213> Conus tulipa

<400> 517  
 His Gly Cys Cys Glu Gly Pro Lys Gly Cys Ser Ser Arg Glu Cys Arg  
 1 5 10 15

Pro Gln His Cys Cys  
 20

<210> 518  
 <211> 23  
 <212> PRT  
 <213> Conus wittigi

<400> 518  
 Leu Pro Ser Cys Cys Asp Phe Glu Arg Leu Cys Val Val Pro Ala Cys  
 1 5 10 15

Ile Arg His Gln Cys Cys Thr  
 20

<210> 519  
 <211> 17  
 <212> PRT  
 <213> Conus omaria

<400> 519  
 Cys Cys Lys Tyr Gly Trp Thr Cys Leu Leu Gly Cys Thr Pro Cys Asp  
 1 5 10 15

Cys

<210> 520

132

<211> 17  
<212> PRT  
<213> Conus omaria

<400> 520  
Cys Cys Arg Tyr Gly Trp Thr Cys Trp Leu Gly Cys Thr Pro Cys Gly  
1 5 10 15

Cys